#### SEQUENCE LISTING

<110> Neo-Morgan Laboratories Inc.
FURUSAWA, Mitsuru

<120> Method and system for rapidly conferring a desired trait to an organism

<130> NE0001PCT

<150> JP 2003-92898

<150> 2003-3-28

<150> US 10/684, 141

<151> 2003-10-10

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<170> Patentin Ver. 2.1

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Gin Met Giu His Asp Met Ala Asp Gin Giu Giu His Asp Leu Ser Ser 85 90 95

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| Lys        | Asp<br>130 | Glu                     | Asn         | Thr        | Ser        | Thr<br>135 | Val        | Val         | Arg        | Phe        | Phe<br>140 | Gly        | Val        | Thr        | Ser        |
| Glu<br>145 | Gly        | His                     | Ser         | Val        | Leu<br>150 | Cys        | Asn        | Val         | Thr        | Gly<br>155 | Phe        | Lys        | Asn        | Tyr        | Leu<br>160 |
| Tyr        | Val        | Pro                     | Ala         | Pro<br>165 | Asn        | Ser        | Ser        | Asp         | Ala<br>170 | Asn        | Asp        | Gln        | Glu        | Gln<br>175 | lle        |
| Asn        | Lys        | Phe                     | Va I<br>180 | His        | Tyr        | Leu        | Asn-       | Glu<br>185  | Thr        | Phe        | Asp        | His        | Ala<br>190 | He         | Asp        |
| Ser        | ile        | Glu<br>195 <sub>.</sub> | Val         | Val        | Ser        | Lys        | GIn<br>200 | Ser         | He         | Trp        | Gly        | Tyr<br>205 | Ser        | Gly        | Asp        |
| Thr        | Lys<br>210 | Leu                     | Pro         | Phe        | Trp        | Lys<br>215 | He         | Tyr         | Val        | Thr        | Tyr<br>220 | Pro        | His        | Met        | Val        |
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| Arg        | Leu        | Met                     | Va I<br>260 | Asp        | Cys        | Gly        | lle        | Va I<br>265 | Gly        | Met        | Ser.       | Trp        | lle<br>270 | Thr        | Leu        |
| Pro        | Lys        | Gly<br>275              | Lys         | Tyr        | Ser        | Met        | lle<br>280 | Glu         | Pro        | Asn        | Asn        | Arg<br>285 | Val        | Ser        | Ser        |
| Cys        | Gln        | Leu                     | Glu         | Val        | Ser        | lle        | Asn        | Tyr         | Arg        | Asn        | Leu        | lle        | Ala        | His        | Pro        |

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| Asp        | lle        | Glu        | Cys         | Ala<br>325                | Gly        | Arg        | He          | Gly        | Va I<br>330 | Phe        | Pro              | Glu        | Pro        | Glu<br>335 | Tyr                |
| Asp        | Pro        | Val        | 11e<br>340  | GIn                       | lle        | Ala        | Asn         | Val<br>345 | Val         | Ser        | He               | Ala        | Gly<br>350 | Ala        | Lys                |
| Lys        | Pro        | Phe<br>355 | He          | Arg                       | Asn        | Val        | Phe<br>360  | Thr        | Leu         | Asn        | Thr              | Cys<br>365 | Ser        | Pro        | lle                |
| Thr        | Gly<br>370 | Ser        | Met         | He                        | Phe        | Ser<br>375 | His         | Ala        | Thr         | Glu        | Glu<br>380       | Glu        | Met        | Leu        | Ser                |
| Asn<br>385 | Trp        | Arg        | Asn         | Phe                       | 11e<br>390 | lle        | Lys         | Vai        | Asp         | Pro<br>395 | Asp              | Val        | lle        | lle        | Gly<br>400         |
| Tyr        | Asn        | Thr        | Thr         | <b>A</b> sn<br><b>405</b> | Phe        | Asp        | He          | Pro        | Tyr<br>410  | Leu        | Leu              | Asn        | Arg        | Ala<br>415 | Lys                |
| Ala        | Leu        | Lys        | Va l<br>420 | Asn                       | Asp        | Phe        | Pro         | Tyr<br>425 | Phe         | Gly        | Arg              | Leu        | Lys<br>430 | Thr        | Val                |
| Lys        | Gln        | Glu<br>435 | He          | Lys                       | Glu        | Ser        | Va I<br>440 | Phe        | Ser         | Ser        | Lys              | Ala<br>445 | Tyr        | Gly        | Thr                |
| Arg        | Glu<br>450 |            | Lys         | Asn                       | Val        | Asn<br>455 | lle         | Asp        | Gly         | Arg        | Leu<br>460       | Gln        | Leu        | Asp        | Leu                |
| Leu<br>465 | Gln        | Phe        | lle         | Gln                       | Arg<br>470 | Glu        | Tyr         | Lys        | Leu         | Arg<br>475 |                  | Tyr        | Thr        | Leu        | <b>A</b> sn<br>480 |
| Ala        | Val        | Ser        | Ala         | His<br>485                | Phe        | Leu        | Gly         | Glu        | Gin<br>490  |            | Glu              | Asp        | Val        | His<br>495 | Tyr                |

| Ser                     | · l le     | ) lle      | 500        | · Asp       | Leu        | Glr         | ı <b>A</b> sn | GI)<br>505 |            | Ser        | · Glu      | ı Thr      | 510        |            | s Ari       |
|-------------------------|------------|------------|------------|-------------|------------|-------------|---------------|------------|------------|------------|------------|------------|------------|------------|-------------|
| Leu                     | Ala        | Val<br>515 |            | · Cys       | Leu        | Lys         | 520           |            | ı Tyr      | · Leu      | Pro        | Leu<br>525 |            | , Leu      | ı Met       |
| Glu                     | Lys<br>530 |            | Met        | : Ala       | Leu        | Va I<br>535 |               | Tyr        | Thr        | Glu        | Met<br>540 |            | Arg        | ∖ Val      | Thr         |
| GIy<br>545              | Val        | Pro        | Phe        | Ser         | Tyr<br>550 | Leu         | Leu           | Ala        | Arg        | Gly<br>555 |            | Gin        | lle        | Lys        | Va I<br>560 |
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| <b>A</b> sn             | Met        | GIn        | Ser<br>580 | Gin         | Ala        | Ser         | Asp           | Asp<br>585 |            | Tyr        | Glu        | Gly        | Ala<br>590 | Thr        | Val         |
| lle                     | Glu        | Pro<br>595 | lle        | Arg         | Gly        | Tyr         | Tyr<br>600    | Asp        | Val        | Pro        | He         | Ala<br>605 | Thr        | Leu        | Asp         |
| Phe                     | Asn<br>610 | Ser        | Leu        | Tyr         | Pro        | Ser<br>615  | He            | Met        | Met        | Ala        | His<br>620 | Asn        | Leu        | Cys        | Tyr         |
| Thr<br>625 <sub>.</sub> | Thr        | Leu        | Cys        | Asn         | Lys<br>630 | Ala         | Thr           | Val        | Glu        | Arg<br>635 | Leu        | Asn        | Leu        | Lys        | 11e<br>640  |
| Asp                     | Glu        | Asp        | Tyr        | Va I<br>645 | lle        | Thr         | Pro           | Asn        | Gly<br>650 | Asp        | Tyr        | Phe        | Vai        | Thr<br>655 | Thr         |
| _ys                     | Arg        | Arg        | Arg<br>660 | Gly         | lle        | Leu         |               | lle<br>665 |            | Leu        | Asp        | Glu        | Leu<br>670 | lle        | Ser         |
| \ia.                    |            | Lys<br>675 | Arg        | Ala         | Lys        |             | Asp<br>680    | Leu        | Arg        | Asp        |            | Lys<br>685 | Asp        | Pro        | Phe         |

| Lys        | Arg<br>690 | Asp          | Val        | Leu        | Asn        | Gly<br>695 | Arg        | Gin        | Leu        | Ala         | 700 <sub>.</sub> | Lys        | He          | Ser        | Ala                |
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| Leu        | Ala        | He           | Ser        | Ser<br>725 | Ser        | Val        | Thr        | Ala        | Tyr<br>730 | Gly         | Arg              | Thr        | Met<br>,    | 11e<br>735 | Leu                |
| Lys        | Thr        | Lys          | Thr<br>740 | Ala        | Val        | GIn        | Glu        | Lys<br>745 | Tyr        | Cys         | lle              | Lys        | Asn<br>750  | Gly        | Tyr                |
| Lys        | His        | Asp<br>755   | Ala        | Val        | Val        | Val        | Tyr<br>760 | Gly        | Asp        | Thr         | Asp              | Ser<br>765 | Val         | Met        | Val                |
| Lys        | Phe<br>770 | Gly          | Thr        | Thr        | Asp        | Leu<br>775 | Lys        | Glu        | Ala        | Met         | Asp<br>780       | Leu        | Gly         | Thr        | Glu                |
| Ala<br>785 | Ala        | Lys          | Tyr        | Val        | Ser<br>790 | Thr        | Leu        | Phe        | Lys        | His<br>795  | Pro              | lle        | Asn         | Leu        | Glu<br>800         |
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| Val        | Met<br>850 |              | Lys        | Val        | Leu        | Lys<br>855 |            | He         | Leu        | ı I le      | Glu<br>860       |            | <b>A</b> sn | Val        | Asp                |
| Gly<br>865 |            | L <b>e</b> u | ı Ala      | Phe        | Val<br>870 |            | Glu        | Thr        | lle        | 875         | Asp              | lle        | Leu         | His        | <b>A</b> sn<br>880 |

| Arg         | Val         | Asp        | ile         | Ser<br>885  | Lys        | Leu         | lle         | lle         | Ser<br>890  | Lys        | Thr         | Leu                  | Ala         | Pro<br>895  | Asn         |
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| Arg         | Glu         | Gly<br>915 | Val         | Gly         | Pro        | Asn         | Va I<br>920 | Gly         | Asp         | Arg        | Val         | Asp<br>925           | Tyr         | Val         | He          |
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| Gln         | Leu         | Gln        | Asn         | Pro<br>965  |            | lle         | Ser         | lle         | Va I<br>970 | Ala        | Pro         | He                   | lle         | Gly<br>975  |             |
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| Asp         | Val         | Arg        | , Asp       | Let<br>1045 |            | Glu         | Lys         | Tyr         | Ser         | · Arg      | Leu         | Trp                  | Thr         | GIn<br>1055 |             |
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PCT/JP2004/004378

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| Şer        | Ala        | GIn<br>35   | Gln        | Leu         | Leu        | Asn        | Ala<br>40  | Ser        | Lys       | He         | Asp              | Asp<br>45  | He         | Asp        | Ser        |
| Met        | Met<br>50  | Gly         | Phe        | Glu         | Arg        | Tyr<br>55  | Val        | Pro        | Pro       | GIn        | Tyr<br>60        | Asn        | Gly        | Arg        | Phe        |
| Asp<br>65  | Ala        | Lys         | Asp        | lle         | Asp<br>70  | Gln        | He         | Pro        | Gly       | Arg<br>75  | Val              | Gly        | Trp        | Leu        | Thr<br>80  |
| Asn        | Met        | His         | Ala        | Thr<br>85   | Leu        | Val        | Ser        | GIn        | Glu<br>90 | Thr        | Leu              | Ser        | Ser        | Gly<br>95  | Ser        |
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| Gly        | lle        | Ser<br>115  | Gly        | Val         | Asp        | Phe        | Tyr<br>120 | Phe        | Leu       | Asp        | Glu              | Glu<br>125 | Gly        | Gly        | Ser        |
| Phe        | Lys<br>130 | Ser         | Thr        | <b>Va</b> ļ | Val        | Tyr<br>135 | Asp        | Pro        | Tyr       | Phe        | Phe<br>140       | lle        | Ala        | Cys        | Asn        |
| Asp<br>145 |            | Ser         | Arg        | Val         | Asn<br>150 |            | Val        | Glu        | Glu       | Leu<br>155 | Val              | Lys        | Lys        | Tyr        | Leu<br>160 |
| Glu        | Ser        | Cys         | Leu        | Lys<br>165  |            | Leu        | Gln        | He         | 11e       |            | Ļys              | Glu        | Asp        | Leu<br>175 |            |
| Met        | Asp        | Asn         | His<br>180 | -           | Leu        | Gly        | Leu        | GIn<br>185 |           | Thr        | Leu              | lle        | Lys<br>190 |            | Ser        |
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| lle         | Leu<br>210 | Gin        | Asp        | Asn        | Ala        | Asn<br>215 | Asn        | Asn        | Val        | Gin         | Arg<br>220 <sub>.</sub> | Asn        | 116        | lyr        | Asn        |
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| Lys         | Asp        | He         | Arg<br>260 | Val        | Gly        | Lys        | Trp        | Tyr<br>265 | Lys        | Val         | Thr                     | Gin        | Gin<br>270 | Gly        | Phe        |
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| Tyr         | Thṛ        | Pro        | Lys<br>340 |            | Glu        | Tyr        | Pro        | Gly<br>345 | Phe        | Phe         | Thr                     | He         | Phe<br>350 | Asn        | Glu        |
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| Pro<br>385  |            | He         | His        | Asn        | Arg<br>390 |            | Lys        | He         | His        | Gly<br>395  |                         | Asp        | Met        | Phe        | Asp<br>400 |

| 3lu        | lle        | Gly        | Phe        | Ala<br>405  | Pro        | Asp        | Ala        | Glu        | Gly<br>410 | Glu        | Tyr        | Lys        | Ser        | Ser<br>415 | Tyr        |
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| He         | Leu<br>530 | •          | Pro        | Asn         | Lys        | His<br>535 |            | Asp        | Pro        | lle        | Glu<br>540 |            | Phe        | Tyr        | Asp        |
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|            |            | 595        |            |            |            |                    | 600        |            |            |            |                         | 605        |             |             |             |
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| Lys<br>625 | Glu        | Asn        | Asn        | He         | Arg<br>630 | <b>A</b> sn        | Glu        | Leu        | Pro        | Leu<br>635 | He                      | Tyr        | His         | Val         | Asp<br>640  |
| Val        | Ala        | Ser        | Met        | Tyr<br>645 | Pro        | Asn                | He         | Met        | Thr<br>650 | Thr        | Asn                     | Arg        | Leu         | GIn<br>655  | Pro         |
| Asp        | Ser        | lle        | Lys<br>660 | Ala        | Glu        | Arg                | Asp        | Cys<br>665 | Ala        | Ser        | Cys                     | Asp        | Phe<br>670  | Asn         | Arg         |
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| Phe        | Phe<br>690 | Pro        | Ser        | Lys        | Met        | <b>A</b> sp<br>695 | Glu        | Tyr        | Asņ        | Met        | lle<br>700              | Lys        | Arg         | Ala         | Leu         |
| GIn<br>705 | Asn        | Glu        | Thr        | Phe        | Pro<br>710 | Asn                | Lys        | Asn        | Lys        | Phe<br>715 |                         | Lys        | Lys         | Lys         | Va I<br>720 |
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| Ser        | Giu        | 11e<br>755 | Val        | Glu        | Arg        | Glu                | Ala<br>760 |            | Val        | Cys        | GIn                     | Arg<br>765 |             | Asn         | Pro         |
| Phe        | Tyr<br>770 |            | Asp        | Thr        | Val        | Lys<br>775         |            | Phe        | Arg        | Asp        | Arg<br>780              |            | Tyr         | Glu         | Phe         |
| Lys<br>785 |            | Leu        | Ala        | Lys        | Thr<br>790 |                    | Lys        | Gly        | Asn        | Leu<br>795 |                         | Lys        | lle         | Asp         | Pro<br>800  |

| Ser         | Asp        | Lys        | His        | Ala<br>805 | Arg        | Asp        | Glu        | Ala        | Lys<br>810 | Lys        | Met        | He         | Val        | Leu<br>815 | Tyr        |
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| Gln                 | Leu        | l le<br>995 | Lys         | Asn         | Phe         |             | Ser<br>1000         | Asp         | lle         | Phe         |             | Val<br>1005 | Phe         | Leu         | Glu         |
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|                     | Asp<br>010 | Thr         | Leu         | Glu         |             | Cys<br>1015 | Tyr                 | Ser         | Ala         |             | Ala<br>1020 | Ser         | Val         | Cys         | Asn         |
| Arg<br>1025         |            | Leu         | Asp         |             | Leu<br>1030 | Asp         | Ser                 | His         |             | Leu<br>1035 | Met         | Leu         | Glu         |             | Glu<br>1040 |
| Asp                 | Leu        | Val         |             | Leu<br>1045 | lle         | Cys         | Glu                 |             | Arg<br>1050 | Ser         | Met         | Ser         |             | Thr<br>1055 | Leu         |
| Lys                 | Glu        | -           | Glu<br>1060 | Gly         | Gln         | Lys         |                     | Thr<br>1065 | Ser         | lle         | Thr         | Thr         | Ala<br>1070 | Arg         | Arg         |
| Leu                 |            | Asp<br>1075 | Phe         | Leu         | Gly         |             | <b>A</b> sp<br>1080 | Met         | Val         | Lys         |             | Lys<br>1085 | Gly         | Leu         | GIn         |
|                     | Lys<br>090 | Tyr         | Ìle         | lle         |             | Ser<br>1095 | Lys                 | Pro         | Phe         |             | Ala<br>1100 | Pro         | Val         | Thr         | Glu         |
| <b>A</b> rg<br>1105 |            | ile         | Pro         |             | Ala<br>1110 |             | Phe                 | Ser         |             | Asp<br>1115 |             | Pro         | ile         |             | Arg<br>1120 |
| Şer                 | Phe        | Leu         |             | Arg<br>1125 |             | Thr         | Leu                 |             | Pro<br>1130 |             | Leu         | Glu         |             | Leu<br>1135 | Asp         |
| lle                 | Arg        |             | lle<br>1140 |             | Asp         | Trp         |                     | Tyr<br>1145 | Tyr         | Arg         | Glu         | Arg         | Leu<br>1150 | Gly         | Ser         |
| Ala                 |            | GIn<br>1155 |             | He          | He          |             | lle<br>1160         | Pro         | Ala         | Ala         |             | GIn<br>1165 |             | Val         | Ser         |
|                     | Pro<br>170 |             | Pro         | Arg         |             | Glu<br>1175 |                     | Pro         | Asp         |             | Leu<br>1180 | Lys         | Arg         | Lys         | lle         |
| Ala                 | Thr        | Lvs         | Glu         | Asn         | Lvs         | Phe         | Lvs                 | Gin         | Thr         | Ser         | Leu         | Thr         | Lvs         | Phe         | Phe         |

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| Ser l       | ∴ys         | Thr         |               | Asn<br>205       | Val         | Pro         | Thr         |             | Gly<br>210  | Lys         | lle <sub>.</sub> | Lys         |              | l le (<br>215 | Glu         |
| Asp l       | Leu         |             | Glu<br>1220   | Pro <sub>.</sub> | Thr         | Val         |             | Glu<br>225  | Asp         | Asn         | Ala              |             | l le<br>230  | Lys           | lle         |
| Ala         |             | Thr<br>1235 | Thr           | Lys              | Lys         |             | Ala<br>1240 | Val         | Ser         | Lys         |                  | Lys<br>1245 | Arg          | <b>A</b> sn   | Gln         |
| Leu<br>1    | Thr<br>250  | Asn         | Glu           | Glu              |             | Pro<br>1255 | Leu         | Val         | Leu         |             | Ser<br>1260      | Glu         | He           | Pro           | Ser         |
| Met /       |             | Glu         | Asp           |                  | Val<br>1270 | Gly         | Trp         | Leu         |             | Tyr<br>1275 | Gln              | Lys         | He           |               | Trp<br>280  |
| Lys         | lle         | Gln         |               | Arg<br>1285      | Asp         | Arg         | Ľys         |             | Arg<br>1290 |             | Gln              | Leu         |              | Gly<br>1295   | Asn         |
| Thr         | Asn         |             | Ser<br>1300   |                  | Glu         | Arg         |             | Ala<br>1305 | Leu         | Gly         | Ser              |             | l le<br>1310 | Arg           | Lys         |
| Gln         |             | Glu<br>1315 |               | Tyr              | Ala         |             | Ser<br>1320 |             | Trp         | Glu         |                  | Leu<br>1325 | Gln          | Tyr           | Lys         |
| -           | Ser<br> 330 |             | Glu           | ) Pro            |             | Val<br>1335 |             | Glu         | Val         |             | Va I<br>1340     |             | lle          | Asn           | Gly         |
| Lys<br>1345 |             | Gln         | <b>A</b> sr   | ı lle            | Thr<br>1350 |             | His         | lle         |             | Lys<br>1355 |                  | lle         | Tyr          | Met           | Lys<br>1360 |
| Phe         | Lys         | Ser         | · Gir         | 1365             |             | : Pro       | Leu         | Gin         | Lys<br>1370 |             | Lys              | Asn         |              | Leu<br>1375   | lle         |
| Glu         | Lys         | Ser         | - Ser<br>1380 |                  | a Ser       | · Leu       | ı Pro       | Asr<br>1385 |             | n Pro       | Lys              |             | Ser<br>1390  | Asn           | Pro         |

| Ala         | -           | GIY<br>1395 | Gin         | Leu         | Pne         |             | 400         | inr         | Leu         | Pro         | 1           | <b>405</b>  | vai         | rie         | Leu         |
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|             | Glu<br>410  | Lys         | Glu         | Asn         |             | Thr<br>415  | Ser         | lle         | Phe         |             | Asp<br> 420 | Glu         | Asn         | Val         | Leu         |
| Gly<br>1425 |             | Phe         | Glu         |             | Thr<br>1430 | lle         | Thr         | Pro         |             | GIn<br>435  | Arg         | Ala         | lle         |             | Asp<br>440  |
| Leu         | Gly         | Ala         |             | Val<br>1445 | Thr         | Phe         | Arg         |             | Lys<br> 450 | Ala         | Met         | Gly         |             | Leu<br>455  | Gly<br>,    |
| Lys         | Gly         |             | GIn<br>1460 | GIn         | Gly         | Phe         |             | Met<br>1465 | Lys         | Asp         | Leu         |             | Met<br>1470 | Ala         | Glu         |
| Asn         |             | Arg<br>1475 |             | Leu         | Ser         |             | Phe<br>1480 | Ser         | Met         | Asp         | lle<br>1    | Gly<br>1485 | Tyr         | Leu         | Lei         |
|             | Phe<br>1490 |             | Thr         | Ser         |             | Gly<br>1495 | Tyr         | Glu         | Phe         |             | Ser<br>1500 | Leu         | Phe         | Lys         | Sei         |
| Trp<br>150  |             | Asp         | Thr         |             | Thr<br>1510 | He          | Leu<br>,    | Val         |             | Lys<br>1515 | Pro         | Ser         | Asn         |             | Ala<br>1520 |
| Gln         | Glu         | lle         |             | Ala<br>1525 |             | Ser         | Leu         |             | GIn<br>1530 | lle         | Tyr         | Lys         |             | Met<br>1535 | Pho         |
| Glu         | Lys         |             | Lys<br>1540 |             | Lys         | He          |             | Thr<br>1545 |             | Ser         | Tyr         |             | Val<br>1550 | Asp         | П           |
| Lys         |             | 1555        |             | Asn         | Phe         |             | Phe<br>1560 |             | Tyr         | Phe         | Thr         | Asp<br>1565 |             | Ser         | Ly:         |
|             | Tyr<br>1570 |             | , Arg       | Leu         |             | GIn<br>1575 |             | Thr         | Thr         |             | Leu<br>1580 |             | Glu         | Glu         | Ar          |

| Gly Leu<br>1585 | Gln         | Phe         |             | Leu<br>590  | Leu         | Leu         | GIn                 |                     | Pro<br>1595 | Phe          | l.le        | Thr         |              | Leu<br>600  |
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| Leu Gly         | Thr         |             | Arg<br>1605 | Leu         | Leu         | Asn         |                     | Met<br>1610         | Pro         | He           | Val         |             | Leu 1<br>615 | Ser         |
| Leu Asn         |             | Val<br>1620 | Leu         | Leu         | Pro         |             | Leu<br>1625         | Asn                 | Trp         | Gln          |             | Thr<br>1630 | Leu          | Leu         |
| Lys Lys         | Leu<br>1635 |             | Asn         | His         |             | Leu<br>1640 | Ser                 | Ser                 | Gly         |              | Trp<br>1645 | He          | Ser          | His         |
| Leu IIe<br>1650 |             | Leu         | Ser         |             | Tyr<br>1655 | Ser         | Asn                 | lle                 |             | l le<br>1660 | Cys         | Asn         | Leu          | Arg         |
| Leu Asp<br>1665 | Ser         | Met         |             | Tyr<br>1670 | He          | lle         | Asp                 |                     | Leu<br>1675 |              | Ala         | Arg         |              | Leu<br>680  |
| Lys Lys         | Glu         |             | lle<br>1685 | Val         | Leu         | Trp         |                     | <b>A</b> sn<br>1690 |             | Lys          | Ala         |             | Leu<br>1695  | Pro         |
| Asp His         |             | Gly<br>1700 |             | Gln         | Asn         |             | Phe<br>1 <b>705</b> |                     | Leu         | Asn          |             | Ser<br>1710 | Trp          | lle         |
| Met Asr         | 1715        |             | Glu         | Phe         |             | Lys<br>1720 |                     | Asn                 | Asn<br>·    |              | Gly<br>1725 |             | Tyr          | Asp         |
| Asn Vai<br>1730 |             | Leu         | Asp         |             | Gly<br>1735 |             | Asp                 | Asn                 |             | Thr<br>1740  |             | Asn         | Thr          | He          |
| Leu Thr<br>1745 | · Ser       | · Ala       |             | lle<br>1750 |             | Asp         | Ala                 |                     | Gly<br>1755 |              | Asp         | Leu         |              | Asn<br>1760 |
| Asn Asr         | Met         |             | Ile<br>1765 |             | Asp         | Lys         |                     | Ala<br>1770         |             | He           | Asn         |             | Pro<br>1775  | Ser         |
| Glu Pho         | . Va!       | Hic         | . Aen       | Δla         | Phe         | Ser         | Asn                 | Asr                 | λla         | Leu          | Asn         | Val         | Leu          | Arg         |

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| 1780                    | )                     | 1785                 |                      | 1790                    |
|-------------------------|-----------------------|----------------------|----------------------|-------------------------|
| Gly Met Leu Lys<br>1795 | Glu Trp Trp           | Asp Glu Al<br>1800   | •                    | Alu Asn Ser Thr<br>305  |
| Ala Asp Leu Leu<br>1810 | ı Val Asn Ser<br>1815 |                      | er Trp Val 6<br>1820 | in Asn Pro Asn          |
| Ala Lys Leu Pha<br>1825 | e Asp Gly Let<br>1830 | ı Leu Arg Ty         | yr His Val H<br>1835 | lis Asn Leu Thr<br>1840 |
| Lys Lys Ala Leu         | ı Leu Gin Let<br>1845 | ı Val Asn Gi<br>185  |                      | Ala Leu Gly Ser<br>1855 |
| Thr lie Val Tyr<br>1860 |                       | g Asn Gin ii<br>1865 | le Leu lie l         | ys Thr Asn Lys<br>1870  |
| Tyr Ser Pro Gli<br>1875 | ı Asn Cys Tyı         | r Ala Tyr Se<br>1880 |                      | Met Met Lys Ala<br>885  |
| Val Arg Thr Asi<br>1890 | n Pro Met Ph          |                      | eu Asp Leu /<br>1900 | Asn ile Lys Arg         |
| Tyr Trp Asp Let<br>1905 | u Leu Ile Tr<br>1910  | p Met Asp Ly         | ys Phe Asn I<br>1915 | Phe Ser Gly Leo<br>1920 |
| Ala Cys lle Gl          | u lle Glu Gl<br>1925  | u Lys Glu As<br>193  |                      | Tyr Thr Ala Val<br>1935 |
| Ser Gin Trp Gi<br>194   |                       | s Phe Leu So<br>1945 | er Pro Ile           | Tyr Gin Pro Git<br>1950 |
| Phe Glu Asp Tr<br>1955  | p Met Met II          | e lle Leu A<br>1960  |                      | Leu Lys Thr Lys<br>965  |
| Gln Ser Tyr Le<br>1970  | u Lys Leu As<br>197   |                      | hr Gin Arg<br>1980   | Pro Thr Gin ile         |

| Val A<br>1985 | sn \       | /al        | Lys         | _           | Gln<br>1990 | Asp         | Lys         | Glu         |              | Ser<br>1995 | Val          | Glu         | Asn         |              | Leu<br>:000 |
|---------------|------------|------------|-------------|-------------|-------------|-------------|-------------|-------------|--------------|-------------|--------------|-------------|-------------|--------------|-------------|
| Asn G         | ly F       | Phe        |             | His<br>2005 | Leu         | Phe         | Ser         |             | Pro<br>2010  | Leu         | Met          | Lys         |             | Va I<br>2015 | Lys         |
| Lys L         | eu F       |            | Lys<br>2020 | Asn         | Gin         | GIn         |             | Phe<br>2025 | lle          | Leu         | Asp          |             | GIn<br>2030 | Tyr          | Glu         |
| Ala A         |            | Tyr<br>035 | Val         | lle         | Pro         |             | Leu<br>2040 | Pro         | Gly          | Ser         |              | Leu<br>2045 | Asn         | Val          | Lys         |
| Asn P         | ro I<br>50 | Leu        | Leu         | Glu         |             | Val<br>2055 | Lys         | Ser         | Leu          |             | His<br>2060  | Val         | Met         | Leu          | Leu         |
| Ser L<br>2065 | ys S       | Ser        | Thr         |             | Leu<br>2070 | Glu         | lle         | Arg         |              | Leu<br>2075 | Arg          | Lys         | Glu         |              | Leu<br>2080 |
| Lys I         | ie i       | Phe        |             | Leu<br>2085 | Arg         | Glu         | Phe         |             | Lys<br>2090  | Val         | Ala          | Glu         |             | Lys<br>2095  | Asp         |
| Pro S         | er i       |            | Ser<br>2100 | Leu         | Val         | Val         |             | Asp<br>2105 |              | Leu         | Cys          |             | Tyr<br>2110 | Cys          | Phe         |
| Phe I         |            | Ser<br>115 | Asp         | He          | Asp         |             | Cys<br>2120 | Lys         | Ala          | Ala         |              | Glu<br>2125 | Ser         | lle          | Phe         |
| Ser 0         | 30         | Val        | Arg         | Cys         |             | Lys<br>2135 | Ala         | Phe         | Asn          |             | Va I<br>2140 | Leu         | Leu         | GIn          | Glu         |
| His L<br>2145 | .eu        | lle        | Gln         |             | Leu<br>2150 |             | Ser         | Asp         |              | Glu<br>2155 |              | Tyr         | Leu         |              | GIn<br>2160 |
| Asp L         | .eu        | Arg        |             | Ser<br>2165 |             | Cys         | His         |             | Va I<br>2170 |             | Arg          | Asp         |             | Met<br>2175  | Ser         |

Ala His Cys Pro Cys Ala Gly Ala Trp Glu Gly Thr Leu Pro Arg Glu

2180

2185

2190

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| cattttcctc | gccaccgcaa  | ggctatcggt | gaccttacag | aaagtattgg | tcttcggaat | 660  |
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| atcgcgctta | agcaacgaaa  | actcgtagtc | cataaagccg | atatcgaacg | ctgcgttatg | 720  |
| gatcaccaac | tccgcgccgc  | gaatatagtc | catgaactca | toggotactt | cggcaaacgt | 780  |
| gggcttatcg | agcaaaaaatt | catcggcaat | accatgtacg | ccaaaggctt | ccggatccac | 840  |
| cagccgatcg | ggtttgagat  | aaacatggaa | gttattgccc | gtcaggcgac | ggttcaccac | 900  |
| ttcaacggca | ccaatctcaa  | tgatcttgtg | gccttcatag | tgcgcaccaa | tctggttcat | 960  |
| accggtggtt | tcggtatcga  | gaacgatctg | gcgtgtaatt | gcagtgctca | tagcggtcat | 1020 |
| ttatgtcaga | cttgtcgttt  | tacagttcga | ttcaattaca | ggaagtctac | cagagatgct | 1080 |
| taaacaggta | gaaattttca  | ccgatggttc | gtgtctgggc | aatccaggac | ctgggggtta | 1140 |
| cggcgctatt | ttacgctatc  | gcggacgcga | gaaaaccttt | agcgctggct | acacccgcac | 1200 |
| caccaacaac | cgtatggagt  | tgatggccgc | tattgtcgcg | ctggaggcgt | taaaagaaca | 1260 |
| ttgcgaagtc | attttgagta  | ccgacagcca | gtatgtccgc | cagggtatca | cccagtggat | 1320 |
| ccataactgg | aaaaaacgtg  | gctggaaaac | cgcagacaaa | aaaccagtaa | aaaatgtcga | 1380 |
| tctctggcaa | cgtcttgatg  | ctgcattggg | gcagcatcaa | atcaaatggg | aatgggttaa | 1440 |
| aggccatgcc | ggacacccgg  | aaaacgaacg | ctgtgatgaa | ctggctcgtg | ccgcggcgat | 1500 |
| gaatcccaca | ctggaagata  | caggctacca | agttgaagtt | taagcctgtg | gtttacgaca | 1560 |
| ttgccgggtg | gctccaaccg  | cctagcgaat | tc         |            | •          | 1592 |
|            |             |            |            |            |            |      |

<210> 42

<211> 243

<212> PRT

<213> Escherichia coli

<400> 42

Met Ser Thr Ala IIe Thr Arg Gin IIe Val Leu Asp Thr Glu Thr Thr 1 5 10 15

Gly Met Asn Gln He Gly Ala His Tyr Glu Gly His Lys He He Glu 20 25 30

ile Gly Ala Val Glu Val Val Asn Arg Arg Leu Thr Gly Asn Asn Phe

His Val Tyr Leu Lys Pro Asp Arg Leu Val Asp Pro Glu Ala Phe Gly 50 55 60

| 65<br>65   | His        | Gly        | He          | Ala        | Asp<br>70  | Glu         | Phe        | Leu        | Leu         | Asp<br>75  | Lys        | Pro        | Thr        | Phe        | A1a<br>80  |
|------------|------------|------------|-------------|------------|------------|-------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|
| Glu        | Val        | Ala        | Asp         | Glu<br>85  | Phe        | Met         | Asp        | Tyr        | 11e<br>90   | Arg        | Gly        | Ala        | Glu        | Leu<br>95  | Val        |
| lle        | His        | Asn        | 'Ala<br>100 | Ala        | Phe        | Asp         | He         | Gly<br>105 | Phe         | Met        | Asp        | Tyr        | Glu<br>110 | Phe        | Ser        |
| Leu        | Leu        | Lys<br>115 | Arg         | Asp        | lle        | Pro         | Lys<br>120 | Thr        | Asn         | Thr        | Phe        | Cys<br>125 | Lys        | Val        | Thr        |
| Asp        | Ser<br>130 | Leu        | Ala         | Val        | Ala        | Arg<br>135  | Lys        | Met        | Phe         | Pro        | Gly<br>140 | Lys        | Arg        | Asn        | Ser        |
| Leu<br>145 | Asp        | Ala        | Leu         | Cys        | Ala<br>150 | Arg         | Tyr        | Glu        | lle         | Asp<br>155 | Asn        | Ser        | Lys        | Arg        | Thr<br>160 |
| Leu        | His        | Gly        | Ala         | Leu<br>165 | Leu        | Asp         | Ala        | Gln        | i le<br>170 |            | Ala        | Glu        | Val        | Tyr<br>175 | Leu        |
| Ala        | Met        | Thr        | Gly<br>180  | Gly        | Gin        | Thr         | Ser        | Met<br>185 |             | Phe        | Ala        | Met        | Glu<br>190 | Gly        | Glu        |
| Thr        | Gln        | GIn<br>195 | Gin         | Gin        | Gly        | Glu         | Ala<br>200 | Thr        | lle         | GIn        | Arg        | 11e<br>205 |            | Arg        | Gin        |
| Ala        | Ser<br>210 |            | Leu         | Arg        | Val        | Va I<br>215 | Phe        | Ala        | Thr         | Asp        | Glu<br>220 |            | lle        | Ala        | Ala        |
| His<br>225 |            | Ala        | Arg         | Leu        | Asp<br>230 |             | Vai        | GIn        | Lys         | Lys<br>235 |            | Gly        | Ser        | Cys        | Leu<br>240 |
| Т          | A          | ۸۱۵        |             |            |            |             |            |            |             |            |            |            |            |            |            |

<210> 43

<211> 4866

<212> DNA

<213> Bacillus subtilis

#### <400> 43

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| tatcaaagat | caagaccgta   | tcatgcaaca | ttacttgccg | tgaacagcac   | gggacttaaa  | 1920 |
|------------|--------------|------------|------------|--------------|-------------|------|
| aatttattta | agcttgtgtc   | actttctcat | attoattatt | tttacagagt   | gccgcgtatt  | 1980 |
| ccgagatctc | agcttgagaa   | atacagggaa | gggcttctga | toggttotgo   | ttgtgacagg  | 2040 |
| ggagaggttt | ttgagggaat   | gatgcaaaaa | tcgcctgaag | aggtggaaga   | tatcgcgtca  | 2100 |
| ttctatgatt | accttgaggt   | tcagccgcct | gaagtgtatc | gtcacttgct   | ggagcttgaa  | 2160 |
| ctggtccgtg | atgaaaaagc   | gctgaaagaa | attattgcga | atatcacgaa   | gctgggggaa  | 2220 |
| aagottaata | aaccggttgt   | tgctacggga | aatgttcatt | acttgaatga   | tgaggataaa  | 2280 |
| atctacagaa | agattttaat   | atcctcacaa | ggcggggcaa | atccgctgaa   | taggcatgaa  | 2340 |
| ctgccgaaag | tgcatttcag   | aacgacagac | gaaatgcttg | aagcttttc    | tttcttaggt  | 2400 |
| gaagaaaaag | cgaaggagat   | cgtagtcacc | aatacccaaa | aggttgcttc   | tttagttgat  | 2460 |
| gacatcaagc | cgattaaaga   | tgatttatat | acgccgaaaa | tcgaaggcgc   | tgatgaagag  | 2520 |
| atcagagaaa | tgagctatca   | gcgtgcaaga | agcatttacg | gggaagagct   | gcctgaaatt  | 2580 |
| gtcgaagcgc | ggattgaaaa   | agagttaaag | agtattattg | gccacggatt   | tgctgttatt  | 2640 |
| tacttgatct | ctcacaaact   | tgtaaaacgt | tcactagatg | acgggtatct   | cgttggttcc  | 2700 |
| cgtggttccg | taggatotto   | attagttgcg | acacttactg | agattactga   | ggtaaacccg  | 2760 |
| ctgccgccgc | actatgtttg   | tcctgagtgc | cagcattctg | agttotttaa   | tgacggttct  | 2820 |
| gtcggttctg | gttttgacct   | gcctgacaag | acatgccctc | attgcggaac   | gcctttgaaa  | 2880 |
| aaagacggcc | atgatattcc   | atttgaaacg | ttcttaggat | ttaaagggga   | caaagtacct  | 2940 |
| gatatcgatt | tgaacttctc   | aggggaatat | cagccgcaag | cacacaatta   | cacaaaagta  | 3000 |
| ttgttcggag | aagacaatgt   | atatcgtgcg | ggaacaatag | gcacggtggc   | agaaaaaaaca | 3060 |
| gcctacggtt | atgtaaaagg   | ctatgccgga | gacaacaatc | ttcatatgcg   | cggtgccgaa  | 3120 |
| atagatcggc | togtacaggg   | atgcacaggt | gtaaaacgta | caactggaca   | gcaccctggc  | 3180 |
| ggtattatcg | tagttccgga   | ttatatggat | atttatgatt | tttcaccgat   | ccagttcccg  | 3240 |
| gcagatgcca | caggttcaga   | gtggaaaacg | actcattttg | atttccactc   | catccatgac  | 3300 |
| aacctgttaa | aacttgatat   | tctcggacac | gatgacccga | ctgttattcg   | gatgcttcaa  | 3360 |
| gacttaagcg | gaatagatco   | gaaaacaatt | ccgacggatg | atcctgaagt   | gatgaagatc  | 3420 |
| ttccagggaa | ccgaatcact   | cggtgtgact | gaagaacaga | ttggctgtaa   | aacgggcact  | 3480 |
|            |              | aacccgattt |            |              |             |      |
|            |              | tcagatttca |            | -            |             |      |
| ggcaatgcad | aggagotcat   | ccacaataat | atttgtgago | tgagtgaggt   | tatcggctgc  | 3660 |
| cgtgatgaca | ttatggttta   | tttaatctat | caaggccttg | agccgtccct   | tgcctttaaa  | 3720 |
| atcatggaat | : tcgtgcgtaa | aggaaaagga | ttaacgcctg | aatgggaaga   | agaaatgaaa  | 3780 |
|            |              | gtatattgat |            |              |             |      |
| aaagcccacg | ccgcggcata   | tgtcttaatg | gcagtccgca | ttgottactt   | taaagtgcat  | 3900 |
| catgotott  | tgtattatgo   | ggcttatttt | accettcete | cagatgactt   | tgatattgat  | 3960 |
| acaatgatca | agggctctac   | agcaatcaga | goggtaatgg | g aggatataaa | cgctaaagga  | 4020 |
| cttgatgctl | caccgaagga   | aaagaacctt | ctgactgttt | tagaattago   | gcttgagatg  | 4080 |

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<210> 44

<211> 1437

<212> PRT

⟨213⟩ Bacillus subtilis

<400> 44

Met Glu Gln Leu Ser Val Asn Arg Arg Gln Phe Gln IIe Leu Leu Gln
1 5 10 15

Gln lle Asn Met Thr Asp Asp Thr Phe Met Thr Tyr Phe Glu His Gly
20 25 30

Glu lle Lys Lys Leu Thr lle His Lys Ala Ser Lys Ser Trp His Phe 35 40 45

His Phe Gin Phe Lys Ser Leu Leu Pro Phe Gin He Tyr Asp Thr Leu 50 55 60

Thr Thr Arg Leu Thr Gln Ser Phe Ala His IIe Ala Lys Val Thr Ser 65 70 75 80

• .

| Ser         | He          | Glu        | Val        | Gin<br>85  | Asp        | Ala        | Glu        | Val         | Ser<br>90  | Glu        | Ser        | ile        | Val        | GIn<br>95  | Asp         |
|-------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|
| Tyr         | Trp         | Ser        | Arg<br>100 | Cys        | He         | Glu        | Glu        | Leu<br>105  | GIn        | Gly        | lle        | Ser        | Pro<br>110 | Pro        | lle         |
| lle         | Ser         | Leu<br>115 | Leu        | Asn        | Gln        | Gin        | Lys<br>120 | Pro         | Lys        | Leu        | Lys        | Gly<br>125 | Asn        | Lys        | Leu         |
| lle         | Va I<br>130 | Lys        | Thr        | Lys        | Thr        | Asp<br>135 | Thr        | Glu         | Ala        | Ala        | Ala<br>140 | Leu        | Lys        | Asn        | Lys         |
| Tyr<br>145  | Ser         | Ser        | Met        | lle        | GIn<br>150 | Ala        | Glu        | Tyr         | Arg        | GIn<br>155 | Phe        | Gly        | Phe        | Pro        | Asp<br>160  |
| Leu         | Gln         | Leu        | Asp        | Ala<br>165 | Glu        | lle        | Phe        | Val         | Ser<br>170 |            | GIn        | Glu        | Val        | Gln<br>175 | Lys         |
| Phe         | Arg         | Glu        | Gln<br>180 |            | Leu        | Ala        | Glu        | Asp<br>185  | Gin        | Glu        | Arg        | Ala        | Met<br>190 |            | Ala         |
| Leu         | lle         | Glu<br>195 |            | Glu        | Lys        | Lys        | Asp<br>200 |             | Glu        | Ser        | Asp        | Glu<br>205 |            | Gln        | Ala         |
| Pro         | Ser<br>210  |            | Pro        | Leu        | Val        | 11e<br>215 |            | Tyr         | Gln        | He         | Lys<br>220 | Asp        | Asn        | Glu        | Glu         |
| l le<br>225 | Arg         | Thr        | Leu        | Asp        | Ser<br>230 |            | Met        | Asp         | Glu        | Glu<br>235 |            | Arg        | lle        | Thr        | Va I<br>240 |
| GIn         | Gly         | Tyr        | Val        | Phe<br>245 |            | Val        | Glu        | Thr         | Arg<br>250 |            | Leu        | Lys        | Ser        | Gly<br>255 | .Arg        |
| Thr         | Leu         | ı Cys      | lle<br>260 |            | Lys        | He         | Thr        | Asp<br>265  |            | Thr        | Asn        | Ser        | 11e<br>270 |            | He          |
| Lys         | Met         | : Phe      | Ala        | Arg        | Glu        | Lys        | Glu        | <b>A</b> sp | Ala        | Ala        | Leu        | Met        | Lys        | Ser        | Leu         |

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|            |            | 275        |            |             |            |             | 280        |            |            |            |                         | 285        |            |             |            |
|------------|------------|------------|------------|-------------|------------|-------------|------------|------------|------------|------------|-------------------------|------------|------------|-------------|------------|
| Lys        | Lys<br>290 | Gly        | Met        | Trp         | Val        | Lys<br>295  | Ala        | Arg        | Gly        | Ser        | 11e <sub>.</sub><br>300 | Gln        | Asn        | Asp         | Thr        |
| Phe<br>305 | Val        | Arg        | Asp        | Leu         | Val<br>310 | Met         | He         | Ala        | Asn        | Asp<br>315 | Val                     | Asn        | Glu        | He          | Lys<br>320 |
| Ala        | Lys        | Thr        | Arg        | Glu<br>325  | Asp        | Ser         | Ala        | Pro        | Glu<br>330 | Gly        | Glu                     | Lys        | Arg        | Va I<br>335 | Glu        |
| Leu        | His        | Leu        | His<br>340 | Ser         | Pro        | Met         | Ser        | GIn<br>345 | Met        | Asp        | Ala                     | Val        | Thr<br>350 | Gly         | He         |
| Gly        | Lys        | Leu<br>355 | Val        | Glu         | GIn        | Ala         | Lys<br>360 | Lys        | Trp        | Gly        | His                     | Glu<br>365 | Ala        | He          | Ala        |
| Leu        | Thr<br>370 |            | His        | Ala         | Val        | Va l<br>375 | Gln        | Ser        | Phe        | Pro        | Asp<br>380              | Ala        | Tyr        | Ser         | Ala        |
| Ala<br>385 | Lys        | Lys        | His        | Gly         | lle<br>390 |             | Met        | He         | Tyr        | Gly<br>395 | Met                     | Glu        | Ala        | Asn         | <b>Leu</b> |
| Val        | Asp        | Asp        | Gly        | Va I<br>405 |            | lle         | Ala        | Tyr        | Asn<br>410 |            | Ala                     | His        | Arg        | Leu<br>415  |            |
| Glu        | Glu        | Glu        | Thr<br>420 |             | Val        | Val         | Phe        | Asp<br>425 |            | Glu        | Thr                     | Thr        | Gly<br>430 |             | Ser        |
| Ala        | Val        | Tyr<br>435 | Asp        | Thr         | lle        | lle         | Glu<br>440 |            | Ala        | Ala        | Val                     | Lys<br>445 |            | Lys         | Gly        |
| Gly        | Glu<br>450 |            | lle        | Asp         | Lys        | Phe<br>455  |            | Ala        | Phe        | Ala        | Asn<br>460              |            | His        | Arg         | Pro        |
| Leu<br>465 |            | Ala        | Thr        | lle         | 11e        |             | Leu        | Thr        | Gly        | 11e<br>475 |                         | Asp        | Asp        | Met         | Le:        |

| GIN        | Asp        | Ala        | Pro        | 485        | vaı        | vai        | Asp        | vai        | 490        | Arg        | ASP .      | rne        | Arg        | 495        | irp        |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| lle        | Gly        | Asp        | Asp<br>500 | He         | Leu        | Val        | Ala        | His<br>505 | Asn        | Ala        | Ser        | Phe        | Asp<br>510 | Met        | Gly        |
| Phe        | Leu        | Asn<br>515 | Val        | Ala        | Tyr        | Lys        | Lys<br>520 | Leu        | Leu        | Glu        | Val        | Glu<br>525 | Lys        | Ala        | Lys        |
| Asn        | Pro<br>530 | Val        | lle        | Asp        | Thr        | Leu<br>535 | Glu        | Leu        | Gly        | Arg        | Phe<br>540 | Leu        | Tyr        | Pro        | Glu        |
| Phe<br>545 | Lys        | Asn        | Hiş        | Arg        | Leu<br>550 | Asn        | Thr        | Leu        | Cys        | Lys<br>555 | Lys        | Phe        | Asp        | lle        | Glu<br>560 |
| Leu        | Thr        | Gln        | His        | His<br>565 | Arg        | Ala        | lle        | Tyr        | Asp<br>570 |            | Glu        | Ala        | Thr        | Ala<br>575 | Tyr        |
| Leu        | Leu        | Leu        | Lys<br>580 |            | Leu        | Lys        | Asp        | Ala<br>585 | Ala        | Glu        | Lys        | Gly        | lle<br>590 | Gln        | Tyr        |
| His        | Asp        | Glu<br>595 |            | Asn        | Glu        | Asn        | Met<br>600 | Gly        | Gln        | Ser        | Asn        | A1a<br>605 |            | Gin        | Arg        |
|            | Arg<br>610 |            | Tyr        | His        | Ala        | Thr<br>615 |            | Leu        | Ala        | Val        | Asn<br>620 |            | Thr        | Gly        | Leu        |
| Lys<br>625 |            | Leu        | Phe        | Lys        | 630        |            | Ser        | Leu        | Ser        | His<br>635 |            | His        | Tyr        | Phe        | Tyr<br>640 |
| Arg        | (Val       | Pro        | Arg        | 645        | : Pro      | Arg        | ; Ser      | GIn        | Leu<br>650 |            | Lys        | Tyr        | Arg        | Glu<br>655 |            |
| Leu        | Leu        | lle        | 660        |            | Ala        | Cys        | . Asp      | Arg<br>665 |            | Glu        | Val        | Phe        | Glu<br>670 |            | Met        |

| Met        | Gln        | Lys<br>675 | Ser            | Pro                        | Glu<br>·            | Glu          | Va I<br>680 | Glu        | Asp        | He         | Ala        | Ser<br>685     | Phe        | Tyr        | Asp         |
|------------|------------|------------|----------------|----------------------------|---------------------|--------------|-------------|------------|------------|------------|------------|----------------|------------|------------|-------------|
| Tyr        | Leu<br>690 | Glu        | Val            | Gin                        | Pro                 | Pro<br>695   | Glu         | Val        | Tyr        | Arg        | His<br>700 | Leu            | Leu        | Glu        | Leu         |
| Glu<br>705 | Leu        | Val        | Arg            | Asp                        | Glu<br>710          | Lys          | Ala         | Leu        | Lys        | Glu<br>715 | He         | He             | Ala        | Asn        | l le<br>720 |
| Thr        | Lys        | Leu        | Gly            | Glu<br>725                 | Lys                 | Leu          | Asn         | Lys        | Pro<br>730 | Val        | Val        | Ala            | Thr        | Gly<br>735 | Asn         |
| Val        | His        | Tyr        | Leu<br>740     | Asn                        | Asp                 | Glu          | Asp         | Lys<br>745 | lle        | Tyr        | Arg        | Lys            | lle<br>750 | Leu        | He          |
| Ser        | Ser        | GIn<br>755 |                | Gly                        | Ala                 | Asn          | Pro<br>760  |            | Asn        | Arg        | His        | Glu<br>765     |            | Pro        | Lys         |
| Val        | His<br>770 |            | Arg            | Thr                        | Thr                 | Asp<br>775   |             | Met        | Leu        | Glu        | Ala<br>780 |                | \$er       | Phe        | Leu         |
| Gly<br>785 |            | Glu        | Lys            | Ala                        | L <u>y</u> s<br>790 |              | He          | Val        | Val        | Thr<br>795 |            | Thr            | Gin        | Lys        | Val<br>800  |
| Ala        | Ser        | Leu        | ı Val          | <b>A</b> sp<br><b>80</b> 5 |                     | lle          | Lys         | Pro        | lle<br>810 |            | Asp        | Asp            | Leu        | Tyr<br>815 | Thr         |
| Pro        | Lys        | ille       | 6 G l u<br>820 |                            | · Ala               | Asp          | Glu         | Glu<br>825 |            | Arg        | G G lu     | Met            | Ser<br>830 |            | Gln         |
| Arg        | , Ala      | Arg<br>835 |                | · He                       | . Tyr               | Gly          | Glu<br>840  |            | r Leu      | Pro        | Glu        | 1   l e<br>845 |            | Glu        | Ala         |
| Are        | 850        |            | ı Lys          | G G I I                    | ı Leu               | 1 Lys<br>855 |             | · He       | lle        | Gly        | / His      |                | Phe        | Ala        | val         |
| He         | . Tyr      | - Lei      | ılle           | e Ser                      | - His               | Lys          | : Leu       | ı Val      | Lys        | Are        | g Ser      | Leu            | ı Asp      | Asp        | Gly         |

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| 865                  |             |            |             |             | 870         |             |             | ٠           |                    | 875         |                     |                     |             |                     | 880         |
|----------------------|-------------|------------|-------------|-------------|-------------|-------------|-------------|-------------|--------------------|-------------|---------------------|---------------------|-------------|---------------------|-------------|
| Tyr                  | Leu         | Val        | Gly         | Ser<br>885  | Arg         | Gly         | Ser         | Val         | Gly<br>890         | Ser         | Ser                 | Leu                 | Val         | Ala<br>895          | Thr         |
| Leu                  | Thr         | Glu        | lle<br>900  | Thr         | Glu         | Val.        | Asn         | Pro<br>905  | Leu                | Pro         | Pro                 | His                 | Tyr<br>910  | Val                 | Cys         |
| Pro                  | Glu         | Cys<br>915 | Gln         | His         | Ser         | Glu         | Phe<br>920  | Phe         | Asn                | Asp         | Gly                 | Ser<br>925          | Val         | Gly                 | Ser         |
| Gly                  | Phe<br>930  | Asp        | Leu         | Pro         | Asp         | Lys<br>935  | Thr         | Cys         | Pro                | His         | Cys<br>940          | Gly                 | Thr         | Pro                 | Leu         |
| Lys<br>945           | Lys         | Asp        | Gly         | His         | Asp<br>950  | He          | Pro         | Phe         | Glu                | Thr<br>955  | Phe                 | Leu                 | Gly         | Phe                 | Lys<br>960  |
| Gly                  | Asp         | Lys        | Val         | Pro<br>965  | Asp         | lle         | Asp         | Leu         | <b>A</b> sn<br>970 | Phe         | Ser                 | Gly                 | Glu         | Tyr<br>975          | GIn         |
| Pro                  | Gln         | Ala        | His<br>980  | Asn         | Tyr         | Thr         | Lys         | Val<br>985  | Leu                | Phe         | Gly                 | Glu                 | Asp<br>990  | Asn                 | Val         |
| Tyr                  | Arg         | Ala<br>995 | Gly         | Thr         | He          |             | Thr<br>1000 | Vai         | Ala                | Glu         |                     | Thr<br>1 <b>005</b> | Ala         | Tyr                 | Gly         |
| _                    | Val<br>1010 | Lys        | Gly         | Tyr         |             | Gly<br>1015 | Asp         | Asn         | Asn                |             | His<br>1 <b>020</b> | Met                 | Arg         | Gly                 | Ala         |
| Glu<br>1 <b>02</b> 5 |             | Asp        | Arg         |             | Val<br>1030 | Gin         | Gly         | Cys         |                    | Gly<br>1035 | Val                 | Lys                 | Arg         | Thr                 | Thr<br>1040 |
| Gly                  | Gin         | His        |             | Gly<br>1045 | Gly         | lle         | lle         |             | Val<br>1050        | Pro         | Asp                 | Tyr                 |             | <b>A</b> sp<br>1055 | lle         |
| Tyr                  | Asp         |            | Ser<br>1060 | Pro         | lle         | Gin         |             | Pro<br>1065 | Ala                | Asp         | Ala                 |                     | Gly<br>1070 | Ser                 | Glu         |

| Trp Lys Thr<br>1075   | Thr His Phe A                | sp Phe His S<br>1080  | Ser lle His As<br>108  | sp Asn Leu Leu<br>35   |
|-----------------------|------------------------------|-----------------------|------------------------|------------------------|
| Lys Leu Asp<br>1090   |                              | lis Asp Asp F<br>1995 | Pro Thr Val I<br>1100  | le Arg Met Leu         |
| Gin Asp Leu :<br>1105 | Ser Gly lle <i>A</i><br>1110 | Asp Pro Lys           | Thr lle Pro Ti<br>1115 | nr Asp Asp Pro<br>1120 |
| Glu Val Met I         | Lys lle Phe 0<br>1125        |                       | Glu Ser Leu G<br>130   | ly Val Thr Glu<br>1135 |
|                       | Gly Cys Lys 1<br>140         | Thr Gly Thr 1<br>1145 | Leu Gly lle P          | ro Glu Phe Gly<br>1150 |
| Thr Arg Phe<br>1155   | Val Arg Gin I                | Met Leu Glu /<br>1160 | Asp Thr Lys P          | ro Thr Thr Phe<br>65   |
| Ser Glu Leu<br>1170   |                              | Ser Gly Leu<br>175    | Ser His Gly T<br>1180  | hr Asp Val Trp         |
| Leu Gly Asn<br>1185   | Ala Gin Giu i<br>1190        | Leu lle His           | Asn Asn Ile C<br>1195  | ys Glu Leu Ser<br>1200 |
| Glu Val IIe           | Gly Cys Arg 1<br>1205        |                       | Met Val Tyr L<br>210   | eu lle Tyr Gin<br>1215 |
|                       | Pro Ser Leu .<br>1220        | Ala Phe Lys<br>1225   | lle Met Glu P          | he Val Arg Lys<br>1230 |
| Gly Lys Gly<br>1235   | Leu Thr Pro                  | Glu Trp Glu<br>1240   |                        | ys Asn Asn Asn<br>145  |
| Val Pro Asp<br>1250   |                              | Asp Ser Cys<br>255    | Lys Lys Ile L<br>1260  | ys Tyr Met Phe         |

| Pro Lys Ala<br>1265 | i His Ala Ala A<br>1270        | Ala Tyr Val Leu                 | Met Ala Val<br>1275   | Arg IIe AIa<br>1280 |
|---------------------|--------------------------------|---------------------------------|-----------------------|---------------------|
| Tyr Phe Lys         | s Val His His <i>I</i><br>1285 | Ala Leu Leu Tyr<br>1290         |                       | Tyr Phe Thr<br>1295 |
| Val Arg Ala         | a Asp Asp Phe /                | Asp lle Asp Thr<br>1305         |                       | Gly Ser Thr<br>310  |
| Ala lle Arg         |                                | Glu Asp IIe Asn<br>1320         | Ala Lys Gly<br>1325   | Leu Asp Ala         |
| Ser Pro Lys         |                                | Leu Leu Thr Val<br>335          | Leu Glu Leu<br>1340   | Ala Leu Glu         |
| Met Cys Glo<br>1345 | ı Arg Gly Tyr :<br>1350        | Ser Phe Gin Lys                 | Val Asp Leu<br>1355   | Tyr Arg Ser<br>1360 |
| Ser Ala Th          | r Glu Phe lle<br>1365          | ile Asp Gly Asr<br>1370         |                       | Pro Pro Phe<br>1375 |
| Asn Ser 11          | e Pro Gly Leu<br>1380          | Gly Thr Asn Ala<br>1385         |                       | lle Val Lys<br>390  |
| Ala Arg Gl<br>139   |                                | Phe Leu Ser Lys<br>1 <b>400</b> | s Glu Asp Leu<br>1405 |                     |
| Gly Lys Va          |                                | lle Leu Glu Tyr<br>415          | Leu Asp Arg           | His Gly Cys         |
|                     |                                |                                 |                       |                     |

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| <b>&lt;212</b> | > PF | RT   |        |      |       |            |      |     |     |      |             |      |     |          |       |
|----------------|------|------|--------|------|-------|------------|------|-----|-----|------|-------------|------|-----|----------|-------|
| <213           | > Ar | abio | lops i | s th | nalia | ana        |      |     |     |      |             |      |     |          |       |
|                |      |      |        | •    |       |            |      |     |     |      |             |      |     |          |       |
| <400           | > 45 | 5    |        |      |       |            |      |     |     |      |             |      |     |          |       |
| Met            | Asn  | Arg  | Ser    | Gly  | He    | Ser        | Lys  | Lys | Arg | Pro  | Pro         | Pro  | Ser | Asn      | Thr   |
| 1              |      |      |        | 5    |       |            |      |     | 10  |      |             |      |     | 15       |       |
|                |      |      |        |      |       |            |      |     |     | -    |             |      |     |          |       |
| Pro            | Pro  | Pro  | Ala    | Gly  | Lys   | His        | Arg  | Ala | Thr | Gly  | Asp         | Ser  | Thr | Pro      | Ser   |
|                |      |      | 20     |      |       |            |      | 25  |     |      |             |      | 30  |          |       |
|                |      |      |        |      |       |            |      |     |     |      |             |      |     |          |       |
| Pro            | Ala  | lle  | Gly    | Thr  | Leu   | Asp        | Asp  | Glu | Phe | Met  | Met         | Glu  | Glu | Asp      | Val   |
|                |      | 35   |        |      |       |            | 40   |     |     |      |             | 45   | •   |          |       |
|                |      |      |        |      |       |            |      |     |     |      |             |      |     | _        |       |
| Phe            |      | Asp  | Glu    | Thr  | Leu   |            | Tyr  | Gly | Asp | Glu  |             | Glu  | Glu | Ser      | Leu   |
|                | 50   |      |        |      |       | 55         |      |     |     |      | 60          |      |     |          |       |
|                |      |      |        |      |       | <b>0</b> 1 |      | ٥.  | ^   |      | <b>.</b>    | •    |     | <b>-</b> | A 1 - |
|                | Leu  | Arg  | Asp    | He   | Glu   | GIU        | Arg  | GIU | Ser |      | Ser         | ser  | Ala | ırp      |       |
| 65             |      |      |        |      | 70    |            |      |     |     | 75   |             |      |     |          | 80    |
| <b>A</b>       | Dura | Dura | Lau    | Com  | Dura  | Ala        | Time | Lau | Cor | Aon  | <b>د</b> م۔ | Gin. | 110 | Dho      | Glo   |
| Arg            | Pro  | Pro  | Leu    |      | Pro   | AIZ        | ıyr  | Leu | 90  | ASII | Ser         | GIII | 116 | 95       | uIII  |
|                |      |      |        | 85   |       |            |      |     | 90  |      |             |      |     | 90       |       |
| Gin            | سم ا | Glu  | ماا    | Aen  | Ser   | ماا        | ماا  | Δla | Glo | Ser  | Hie         | l ve | Glu | Len      | Len   |
| um             | LCU  | uiu  | 100    | nap  | JCI   | 110        | 116  | 105 | uiu | OCI  | 1110        | Lyo  | 110 | Lou      | LCu   |
|                |      |      | 100    |      |       |            |      | 100 |     |      |             |      | 110 |          |       |
| Pro            | GIV  | Ser  | Ser    | Glv  | Gln   | Ala        | Pro  | He  | He  | Arg  | Met.        | Phe  | Glv | Val      | Thr   |
| .,,            | 41,  | 115  | OO.    | 4.,  | ٠     | ,,,,       | 120  |     |     |      |             | 125  | ,   |          |       |
|                |      |      |        |      |       |            | 0    |     |     |      |             |      |     |          |       |
| Arg            | Glu  | Glv  | Asn    | Ser  | Val   | Cvs        | Cvs  | Phe | Val | His  | Gly         | Phe  | Glu | Pro      | Tyr   |
|                | 130  | ,    |        |      |       | 135        |      |     |     |      | 140         |      |     |          | -     |
|                |      |      |        |      |       |            |      |     |     |      | •           |      |     | •        |       |
| Phe            | Tyr  | He   | Ala    | Cys  | Pro   | Pro        | Gly  | Met | Gly | Pro  | Asp         | Asp  | He  | Ser      | Asn   |
| 145            |      |      |        | •    | 150   |            | -    |     | _   | 155  |             |      |     |          | 160   |
|                |      |      |        |      |       |            |      |     |     |      |             |      |     |          |       |
| Phe            | His  | GIn  | Ser    | Leu  | Glu   | Gly        | Arg  | Met | Arg | Glu  | Ser         | Asn  | Lys | Asn      | Ala   |
|                |      |      |        | 165  |       | Ĭ          | -    |     | 170 |      |             |      | •   | 175      |       |
|                |      |      |        |      |       |            |      |     |     |      |             |      |     |          |       |

| Lys        | Val         | Pro        | Lys<br>180 | Phe        | Val        | Lys         | Arg         | ! le<br>185 | Glu        | Met        | Val        | GIn        | Lys<br>190 | Arg         | Ser        |
|------------|-------------|------------|------------|------------|------------|-------------|-------------|-------------|------------|------------|------------|------------|------------|-------------|------------|
| lle        | Met         | Tyr<br>195 | Tyr        | GIn        | Gln        | GIn         | Lys<br>200  | Ser         | Gln        | Thr        | Phe        | Leu<br>205 | Lys        | He          | Thr        |
| Val        | Ala<br>210  | Leu        | Pro        | Thr        | Met        | Va I<br>215 | Ala         | Ser         | Cys        | Arg        | Gly<br>220 | lle        | Leu        | Asp         | Arg        |
| Gly<br>225 | Leu         | GIn        | He         | Asp        | Gly<br>230 | Leu         | Gly         | Met         | Lys        | Ser<br>235 | Phe        | Gln        | Thr        | Tyr         | Glu<br>240 |
| Ser        | Asn         | lle        | Leu        | Phe<br>245 | Val        | Leu         | Arg         | Phe         | Met<br>250 | Val        | Asp        | Cys        | Asp        | lle<br>255  | Val        |
| Gly        | Gly         | Asn        | Trp<br>260 | lle        | Glu        | Val         | Pro         | Thr<br>265  | Gly        | Lys        | Tyr        | Lys        | Lys<br>270 | Asn         | Ala        |
| Arg        | Thr         | Leu<br>275 | Ser        | Tyr        | Cys        | Gln         | Leu<br>280  | Glu         | Phe        | His        | Cys        | Leu<br>285 | Tyr        | Ser         | Asp        |
| Leu        | l le<br>290 | Ser        | His        | Ala        | Alą        | Glu<br>295  | Gly         | Glu         | Tyr        | Ser        | Lys<br>300 | Met        | Ala        | Pro         | Phe        |
| Arg<br>305 | Val         | Leu        | Ser        | Phe        | Asp<br>310 | lle         | Glu         | Cys         | Ala        | Gly<br>315 | Arg        | Lys        | Gly        | His         | Phe<br>320 |
| Pro        | Glu         | Ala        | Lys        | His<br>325 | Asp        | Pro         | Val         | lle         | GIn<br>330 | lle        | Ala        | Asn        | Leu        | Va I<br>335 | Thr        |
| Leu        | Gin         | Gly        | Glu<br>340 | Asp        | His        | Pro         | Phe         | Va i<br>345 | Arg        | Asn        | Vai        | Met        | Thr<br>350 | Leu         | Lys        |
| Ser        | Cys         | Ala<br>355 |            | He         | Val        | Gly         | Va I<br>360 | Asp         | Val        | Met        | Ser        | Phe<br>365 | Gľu        | Thr         | Glu        |

| Arg        | Glu<br>370 | Val        | Leu        | Leu        | Ala        | Trp<br>375  | Arg        | Asp         | Leu        | lle        | Arg<br>380 | Asp         | Val        | Asp        | Pro        |
|------------|------------|------------|------------|------------|------------|-------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|
| Asp<br>385 | lle        | lle        | lle        | Gly        | Tyr<br>390 | Asn         | He         | Cys         | Lys        | Phe<br>395 | Asp        | Leu         | Pro        |            | Leu<br>400 |
| lle        | Glu        | Arg        | Ala        | A1a<br>405 | Thr        | Leu         | Gly        | He          | Glu<br>410 | Glu        | Phe        | Pro         | Leu        | Leu<br>415 | Gly        |
| Arg        | Val        | Lys        | Asn<br>420 | Ser        | Arg        | Val         | Arg        | Va I<br>425 | Arg        | Asp        | Ser        | Thr         | Phe<br>430 | Ser        | Ser        |
| Arg        | Gln        | Gln<br>435 | Gly        | He         | Arg        | Glu         | Ser<br>440 | Lys         | Glu        | Thr        | Thr        | l le<br>445 | Glu        | Gly        | Arg        |
| Phe        | GIn<br>450 | Phe        | Asp        | Leu        | lle        | GIn<br>455  | Ala        | lle         | His        | Arg        | Asp<br>460 | His         | Lys        | Leu        | Ser        |
| Ser<br>465 | Tyr        | Ser        | Leu        | Asn        | Ser<br>470 | Val         | Ser        | Ala         | His        | Phe<br>475 | Leu        | Ser         | Ġlu        | GIn        | Lys<br>480 |
| Glu        | Asp        | Val        | His        | His<br>485 | Ser        | lle         | He         | Thr         | Asp<br>490 | Leu        | Gln        | Asn         | Gly        | Asn<br>495 | Ala        |
| Glu        | Thr        | Arg        | Arg<br>500 | Arg        | Leu        | Ala         | Vai        | Tyr<br>505  | (          | Leu        | Lys        | Asp         | Ala<br>510 | Tyr        | Leu        |
| Pro        | Gln        | Arg<br>515 | Leu        | Leu        | Asp        | Lys         | Leu<br>520 | Met         | Phe        | lie        | Tyr        | Asn<br>525  | Tyr        | Val        | Glu        |
| Met        | Ala<br>530 |            | Val        | Thr        | Gly        | Va I<br>535 |            | lie         | Ser        | Phe        | Leu<br>540 |             | Ala        | Arg        | Gly        |
| GIn<br>545 |            | He         | Lys        | Val        | Leu<br>550 |             | Gln        | Leu         | Leu        | Arg<br>555 |            | Gly         | Lys        | Gin        | Lys<br>560 |
| Asn        | Leu        | Val        | Leu        | Pro        | Asn        | Ala         | Lys        | Gin         | Ser        | Gly        | Ser        | Glu         | Gln        | Gly        | Thr        |

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|   |            |            |            |            | 565        |            |            |            |            | 570        |            |                  |            |            | 575        |            |
|---|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|
| T | yr         | Glu        | Gly        | Ala<br>580 | Thr        | Val        | Leu        | Glu        | Ala<br>585 | Arg        | Thr        | Gly <sub>.</sub> | Phe        | Tyr<br>590 | Glu        | Lys        |
| F | ro         | lle        | Ala<br>595 | Thr        | Leu        | Asp        | Phe        | Ala<br>600 | Ser        | Leu        | Tyr        | Pro              | Ser<br>605 | He         | Met        | Met        |
| F | lla        | Tyr<br>610 | Asn        | Leu        | Cys        | Tyr        | Cys<br>615 | Thr        | Leu        | Val        | Thr        | Pro<br>620       | Glu        | Asp        | Val        | Arg        |
|   | .ys<br>625 | Leu        | Asn        | Leu        | Pro        | Pro<br>630 | Glu        | His        | Val        | Thr        | Lys<br>635 | Thr              | Pro        | Ser        | Gly        | Glu<br>640 |
| 1 | Thr        | Phe        | Val        | Lys        | GIn<br>645 | Thr        | Leu        | Gln        | Lys        | Gly<br>650 | He         | Leu              | Pro        | Glu        | lle<br>655 | Leu        |
| ( | ilu        | Glu        | Leu        | Leu<br>660 | Thr        | Ala        | Arg        | Lys        | Arg<br>665 | Ala        | Lys        | Ala              | Asp        | Leu<br>670 | Lys        | Glu        |
| 1 | λla        | Lys        | Asp<br>675 | Pro        | Leu        | Glu        | Lys        | A1a<br>680 | Val        | Leu        | Asp        | Gly              | Arg<br>685 | GIn        | Leu        | Ala        |
| l | _eu        | Lys<br>690 | He         | Ser        | Ala        | Asn        | Ser<br>695 | Val        | Tyr        | Gly        | Phe        | Thr<br>700       | Gly        | Ala        | Thr        | Vai        |
|   | 31y<br>705 | Gln        | Leu        | Pro        | Cys        | Leu<br>710 | Glu        | lle        | Ser        | Ser        | Ser<br>715 | Val              | Thr        | Ser        | Tyr        | Gly<br>720 |
| 1 | Arg        | Gin        | Met        | lle        | Glu<br>725 | Gin        | Thr        | Lys        | Lys        | Leu<br>730 | Val        | Glu              | Asp        | Lys        | Phe<br>735 | Thr        |
| - | Thr        | Leu        | Gly        | Gly<br>740 |            | Gln        | Tyr        | Asn        | Ala<br>745 |            | Val        | lle              | Tyr        | Gly<br>750 | Asp        | Thr        |
| 1 | Asp        | Ser        | Vai        | Met        | Val        | Gin        | Phe        | Glv        | Val        | Ser        | Asp        | Val              | Glu        | Ala        | Ala        | Met        |

|            |            | 755         |            |            |            |            | 760        |            |            |            |             | 765        |            | -          |            |
|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|
|            | Leu<br>770 | Gly         | Arg        | Glu        | Ala        | Ala<br>775 | Glu        | His        | lle        | Ser        | Gly<br>780  | Thr        | Phe        | He         | Lys        |
| Pro<br>785 | lle        | Lys         | Leu        | Glu        | Phe<br>790 | Glu        | Lys        | Vai        | Tyr        | Phe<br>795 | Pro         | Tyr        | Leu        | Leu        | 11e<br>800 |
| Asn        | Lys        | Lys         | Arg        | Tyr<br>805 | Ala        | Gly        | Leu        | Leu        | Trp<br>810 | Thr        | Asn         | Pro        | GIn        | Gln<br>815 | Phe        |
| Asp        | Lys        | Met         | Asp<br>820 | Thr        | Lys        | Gly        | lle        | Glu<br>825 | Thr        | Val        | Arg         | Arg        | Asp<br>830 | Asn        | Cys        |
| Leu        | Leu        | Va I<br>835 | Lys        | Asn        | Leu        | Val        | Thr<br>840 | Glu        | Ser        | Leu        | Asn         | Lys<br>845 | lle        | Leu        | lle        |
| Asp        | Arg<br>850 | Asp         | Val        | Pro        | Gly        | Ala<br>855 | Ala        | Glu        | Asn        | Val        | Lys<br>860  | Lys        | Thr        | lle        | Ser        |
| Asp<br>865 | Leu        | Leu         | Met        | Asn        | Arg<br>870 | He         | Asp        | Leu        | Ser        | Leu<br>875 | Leu         | Val        | lle        | Thr        | Lys<br>880 |
| Gly        | Leu        | Thr         | Lys        | Thr<br>885 |            | Asp        | Asp        | Tyr        | Glu<br>890 |            | Lys         | Ser        | Ala        | His<br>895 |            |
| Glu        | Leu        | Ala         | Glu<br>900 |            | Met        | Arg        | Lys        | Arg<br>905 | Asp        | Ala        | Ala         | Thr        | Ala<br>910 |            | Asn        |
| Val        | Gly        | Asp<br>915  |            | Val        | Pro        | Tyr        | Val<br>920 |            | lie        | Lys        | Ala         | Ala<br>925 |            | Gly        | Ala        |
| Lys        | Ala<br>930 |             | · Glu      | Arg        | Ser        | Glu<br>935 |            | Pro        | He         | Tyr        | Va I<br>940 |            | Gin        | Asn        | Asr        |
| 11e<br>945 |            | lle         | Asp        | Pro        | Asn<br>950 |            | Tyr        | Leu        | Glu        | Asn<br>955 | GIn         | He         | Ser        | Lys        | Pro<br>960 |

| Leu Leu  | Arg                                     | He                    | Phe                     | Glu         | Pro               | Val         | Leu          | Lys       | Asn        | Ala      | Ser  | Lys         | Glu        | Leu         |
|--|---|-----------------------|-------------------------|-------------|-------------------|-------------|--------------|-----------|------------|----------|------|-------------|------------|-------------|
|  |   |                       | 965                     |             |                   |             |              | 970       |            |          |      |             | 975        |             |
|  |   |                       |                         |             |                   |             |              |           |            |          |      |             |            |             |
| Leu His  |   |                       | His                     | Thr         | Arg               | Ser         |              | Ser       | He         | Thr      | Thr  |             | Ser        | Asn         |
|  |   | 980                   |                         |             |                   |             | 985          |           |            |          |      | 990         |            |             |
| One Olsa   | 11.                                     | ll air                | Lva                     | Dho         | A I o             | Lva         | Lvo          | G)n       | 1 011      | Sor      | ٥,,, | Va I        | Gly :      | Cve         |
| Ser Gly  |   | WEL                   | LyS                     | rile        |                   | Lys<br>1000 | Lys          | GIII      | LEU        |          | 005  | Vai         | ui y       | Uys         |
|  | 995                                     |                       |                         |             |                   | 1000        |              |           |            | ,        | 000  |             |            |             |
| Lys Val  | Pro                                     | ile                   | Arg                     | Tvr         | Phe               | Val         | Gin          | Trp       | Asn        | Thr      | Met  | Arg         | Lys        | Leu         |
| 1010   |   |                       |                         |             | 1015              |             |              |           |            | 1020     |      | Ī           | -          |             |
| ,  |   |                       |                         |             |                   |             |              |           |            |          |      |             |            |             |
| Gln Gly  | Lys                                     | Arg                   | Ser                     | Arg         | Val               | He          | Leu          | GIn       | Lys        | Arg      | Val  | Ser         | Arg        | Tyr         |
| 1025   |   |                       | -                       | 1030        |                   |             |              |           | 1035       |          |      |             | 1          | 040         |
|  |   |                       |                         |             |                   |             |              |           |            |          |      |             |            |             |
| Ala Ala  | Trp                                     | Leu                   | Ser                     | Leu         | Lys               | Arg         | Phe          | Leu       | Gly        | Gly      | Cys  | Gly         | His        | Ser         |
|  |   |                       | 1045                    |             | •                 |             |              | 1050      |            |          |      | •           | 1055       |             |
|  |   |                       |                         |             |                   |             |              |           |            |          |      |             | •          |             |
|  |   |                       |                         |             |                   |             |              |           |            |          |      |             |            |             |
| Ala Arg  | Ser                                     | Val                   | Lys                     | Ala         | Leu               | Phe         | l le         | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Ala Arg  |   | Va I<br>1060          |                         | Ala         | Leu               |             | l le<br>1065 | Lys       | Met        | Ser      |      | Ala<br>1070 | Pro        | Val         |
|  | 1                                       | 1060                  |                         |             | •                 |             | 1065         | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu lle  | Va l                                    | 1060                  |                         |             | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu lle  | 1                                       | 1060                  |                         |             | Thr               |             | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu lle  | Va l                                    | 1060                  |                         |             | Thr               | Gly         | 1065<br>Glu  | Lys       | <b>Met</b> | Ser      |      |             | Pro        | Val         |
| Glu I le   | Val<br>1075                             | 1060                  |                         |             | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu lle<br><210≻ 4   | Val<br>1075<br>6                        | 1060                  |                         |             | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | <b>V</b> al |
| Glu I le<br><210> 4<br><211> 2   | Val<br>1075<br>6<br>154                 | 1060                  |                         |             | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu l le  <210> 4  <211> 2  <212> P  | Val<br>1075<br>6<br>154<br>RT           | 1060<br>GIn           | Tyr                     | Phe         | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu I le<br><210> 4<br><211> 2   | Val<br>1075<br>6<br>154<br>RT           | 1060<br>GIn           | Tyr                     | Phe         | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser      |      |             | Pro        | Val         |
| Glu l le  <210> 4  <211> 2  <212> P  | Val<br>1075<br>6<br>154<br>RT<br>rabio  | 1060<br>GIn           | Tyr                     | Phe         | Thr               | Gly         | 1065<br>Glu  | Lys       | Met        | Ser .    |      |             | Pro        | Val         |
| Glu I le  <210> 4  <211> 2  <212> P  <213> A   | Val<br>1075<br>6<br>154<br>RT<br>rabid  | GIn                   | Tyr                     | Phe<br>hali | Thr               | Gly<br>1080 | <b>Glu</b>   |           |            |          |      | 1070        |            |             |
| Glu I le <210> 4 <211> 2 <212> P <213> A <400> 4   | Val<br>1075<br>6<br>154<br>RT<br>rabid  | GIn                   | Tyr                     | Phe<br>hali | Thr               | Gly<br>1080 | <b>Glu</b>   |           | Asn        |          |      | 1070        |            |             |
| <pre>&lt;210&gt; 4 &lt;210&gt; 4 &lt;211&gt; 2 &lt;212&gt; P &lt;213&gt; A &lt;400&gt; 4 Met Ser</pre> | Val<br>1075<br>6<br>154<br>RT<br>rabid  | GIn                   | Tyr<br>is t             | Phe<br>hali | Thr               | Gly<br>1080 | <b>Glu</b>   | Leu       | Asn        |          |      | 1070        | Val        |             |
| <pre>&lt;210&gt; 4 &lt;210&gt; 4 &lt;211&gt; 2 &lt;212&gt; P &lt;213&gt; A &lt;400&gt; 4 Met Ser</pre> | Val<br>1075<br>6<br>1154<br>RT<br>rabio | (1060)<br>Gln<br>dops | Tyr<br>is t<br>Arg<br>5 | Phe hali    | Thr<br>ana<br>Asp | Gly<br>1080 | Glu          | Leu<br>10 | Asn        | ·<br>Val | Gln  | 1070<br>Lys | Va I<br>15 | Ser         |

| Gly        | Glu        | Thr<br>35  | Arg        | Leu        | Gly        | Trp        | Leu<br>40  | Leu         | Thr        | Phe        | Ala         | Ser<br>45  | Ser        | Ser        | Trp        |
|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|-------------|------------|------------|------------|------------|
| Glu        | Asp<br>50  | Ala        | Asp        | Thr        | Gly        | Lys<br>55  | Thr        | Phe         | Ser        | Cys        | Va I<br>60  | Asp        | Leu        | Phe        | Phe        |
| Va I<br>65 | Thr        | Gln        | Asp        | Gly        | Ser<br>70  | Ser        | Phe        | Lys         | Thr        | Lys<br>75  | Tyr         | Lys        | Phe        | Arg        | Pro<br>80  |
| Tyr        | Leu        | Tyr        | Ala        | Ala<br>85  | Thr        | Lys        | Asp        | Asn         | Met<br>90  | Glu        | Leu         | Glu        | Val        | Glu<br>95  | Ala        |
| Tyr        | Leu        | Arg        | Arg<br>100 | Arg        | Tyr        | Glu        | Arg        | GIn<br>105  | Val        | Ala        | Asp         | lle        | Gin<br>110 | lle        | Val        |
| His        | Lys        | Glu<br>115 | Asp        | Leu        | Tyr        | Leu        | Lys<br>120 | Asn         | His        | Leu        | Ser         | Gly<br>125 | Leu        | Gln        | Lys        |
| Lys        | Tyr<br>130 | Leu        | Lys        | Val        | Ser        | Phe<br>135 |            | Thr         | Val        | Gin        | Gin<br>140  | Leu        | Val        | Glu        | Val        |
| Lys<br>145 |            | Asp        | Leu        | Leu        | His<br>150 | He         | Val        | Glu         | Arg        | Asn<br>155 |             | Ala        | Lys        | Phe        | Asn<br>160 |
| Ala        | Leu        | Glu        | Ala        | Tyr<br>165 | Glu        | Ser        | He         | Leu         | Ser<br>170 |            | Lys         | Arg        | Glu        | GIn<br>175 | Arg        |
| Pro        | GIn        | Asp        | Cys<br>180 |            | Asp        | Ser        | Val        | Va l<br>185 |            | Leu        | Arg         | Glu        | Tyr<br>190 | Asp        | Val        |
| Pro        | Tyr        | His<br>195 |            | Arg        | Phe        | Ala        | lle<br>200 |             | Asn        | Asp        | Val         | Arg<br>205 |            | Gly        | Gin        |
| Trp        | Tyr<br>210 |            | Val        | Ser        | lle        | Ser<br>215 |            | Thr         | Asp        | Val        | l le<br>220 |            | Glu        | Lys        | Arg        |

| Thr<br>225  | Asp        | Leu        | Leu        | Gln        | Arg<br>230 | Ala        | Glu        | Val        | Arg         | Va I<br>235 | Cys        | Ala        | Phe        | Asp        | ile<br>240 |
|-------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|-------------|------------|------------|------------|------------|------------|
| Glu         | Thr        | Val        | Lys        | Leu<br>245 | Pro        | Leu        | Lys        |            | Pro<br>250  | Asp         | Ala        | Glu        | Tyr        | Asp<br>255 | Gln        |
| He          | Met        | Met        | lle<br>260 | Ser        | Tyr        | Met        | Val        | Asp<br>265 | Gly         | GIn         | Gly        | Phe        | Leu<br>270 | lle        | Thr        |
| <b>A</b> sn | Arg        | Glu<br>275 | Cys        | Val        | Gly        | Lys        | Asp<br>280 | lle        | <b>Gl</b> u | Asp         | Leu        | Glu<br>285 | Tyr        | Thr        | Pro        |
| Lys         | Pro<br>290 | Glu        | Phe        | Glu        | Gly        | Tyr<br>295 | Phe        | Lys        | Val         | Thr         | Asn<br>300 | Val        | Thr        | Asn        | Glu        |
| Val<br>305  | Glu        | Leu        | Leu        | Arg        | Lys<br>310 | Trp        | Phe        | Ser        | His         | Met<br>315  | Gln        | Glu        | Leu        | Lys        | Pro<br>320 |
| Gly         | lle        | Tyr        | Val        | Thr<br>325 | Tyr        | Asn        | Gly        | Asp        | Phe<br>330  | Phe         | Asp        | Trp        | Pro        | Phe<br>335 | lle        |
| Glu         | Arg        | Arg        | Ala        | Ser        | His        | His        | Gly        | lle        | Lys         | Met         | Asn        | Glu        | Glu        | Leu        | Gly        |
|             |            |            | 340        |            |            |            |            | 345        |             |             |            |            | 350        |            |            |
| Phe<br>+    | Arg        | Cys<br>355 |            | Gin        | Asn        | Gln        | Gly<br>360 |            | Cys         | Arg         | Ala        | Lys<br>365 |            | Val        | Cys        |
| His         | Leu<br>370 |            | Cys        | Phe        | Ser        | Trp<br>375 |            | Lys        | Arg         | Asp         | Ser<br>380 | Tyr        | Leu        | Pro        | Gln        |
| Gly<br>385  | Ser        | GIn        | Gly        | Leu        | Lys<br>390 |            | Val        | Thr        | Lys         | Va I<br>395 |            | Leu        | Gly        | Tyr        | Asp<br>400 |
| Pro         | Leu        | Glu        | Val        | Asn<br>405 | Pro        | Glu        | Asp        | Met        | Val<br>410  |             | Phe        | Ala        | Met        | Glu<br>415 | Lys        |

| Pro          | GIn        | Thr         | Met<br>420 | Ala        | Ser        | Tyr        | Ser         | Va I<br>425 | Ser        | Asp        | Ala        | Val        | Ala<br>430 | Thr        | Tyr        |
|--------------|------------|-------------|------------|------------|------------|------------|-------------|-------------|------------|------------|------------|------------|------------|------------|------------|
| Tyr          | Leu        | Tyr<br>435  | Met        | Thr        | Tyr        | Val        | His<br>440  | Pro         | Phe        | Val        | Phe        | Ser<br>445 | Leu        | Ala        | Thr        |
| lle          | lle<br>450 | Pro         | Met        | Val        | Pro        | Asp<br>455 | Glu         | Val         | Leu        | Arg        | Lys<br>460 | Gly        | Ser        | Gly        | Thr        |
| Leu<br>465   | Cys        | Glu         | Met        | Leu        | Leu<br>470 | Met        | Val         | Glu         | Ala        | Tyr<br>475 | Lys        | Ala        | Asn        | Val        | Val<br>480 |
| Cys          | Pro        | Asn         | Lys        | Asn<br>485 | Gl'n       | Ala        | Asp         | Pro         | Glu<br>490 | Lys        | Phe        | Tyr        | GIn        | Gly<br>495 | Lys        |
| Leu          | Leu        | Glu         | Ser<br>500 | Glu        | Thr        | Tyr        | lle         | Gly<br>505  | Gly        | His        | Val        | Glu        | Cys<br>510 | Leu        | GIn        |
| Ser          | Gly        | Va I<br>515 | Phe        | Arg        | Ser        | Asp        | l le<br>520 | Pro         | Thr        | Ser        | Phe        | Lys<br>525 | Leu        | Asp        | Ala        |
| Ser          | Ala<br>530 | Tyr         | GIn        | Gin        | Leu        | lle<br>535 | Asp         | Asn         | Leu        | Gly        | Arg<br>540 | Asp        | Leu        | Giu        | Tyr        |
| A I a<br>545 | lle        | Thr         | Val        | Glu        | Gly<br>550 | Lys        | Met         | Arg         | Met        | Asp<br>555 | Ser        | Val        | Ser        | Asn        | Phe<br>560 |
| <b>A</b> sp  | Glu        | Val         | Lys        | Glu<br>565 | Val        | He         | Arg         | Glu         | Lys<br>570 | Leu        | Glu        | Lys        | Leu        | Arg<br>575 | Asp        |
| Asp          | Pro        | He          | Arg<br>580 | Glu        | Glu        | Gly        | Pro         | Leu<br>585  | lie        | Tyr        | His        | Leu        | Asp<br>590 | Val        | Ala        |
| Ala          | Met        | Tyr<br>595  | Pro        | Asn        | He         | lle        | Leu<br>600  | Thr         | Asn        | Arg        | Leu        | GIn<br>605 | Pro        | Pro        | Ser        |
| lle          | Val        | Thr         | Asp        | Glu        | Val        | Cys        | Thr         | Ala         | Cys        | Asp        | Phe        | Asn        | Gly        | Pro        | Glu        |

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|            | 610        |            |            |            |            | 615        |            |            |            |            | 620              |            |            |             |          |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|-------------|----------|
| Lys<br>625 | Thr        | Cys        | Leu        | Arg        | Lys<br>630 | Leu        | Glu        | Trp        | Val        | Trp<br>635 | Arg <sub>.</sub> | Gly        | Val        | Thr         | Ph<br>64 |
| Lys        | Gly        | Asn        | Lys        | Ser<br>645 | Glu        | Tyr        | Tyr        | His        | Leu<br>650 | Lys        | Lys              | Gln        | lle        | Glu<br>655  | Se       |
| Glu        | Ser        | Val        | Asp<br>660 | Ala        | Gly        | Ala        | Asn        | Met<br>665 | Gln        | Ser        | Ser              | Lys        | Pro<br>670 | Phe         | Le       |
| Asp        | Leu        | Pro<br>675 | Lys        | Val        | Glu        | Gln        | GIn<br>680 | Ser        | Lys        | Leu        | Lys              | Glu<br>685 | Arg        | Leu         | Lу       |
| Lys        | Tyr<br>690 | Cys        | Gln        | Lys        | Ala        | Tyr<br>695 | Ser        | Arg        | Val        | Leu        | Asp<br>700       | Lys        | Pro        | He          | Th       |
| Glu<br>705 | Val        | Arg        | Glu        | Ala        | Gly<br>710 | lle        | Cys        | Met        | Arg        | Glu<br>715 | Asn              | Pro        | Phe        | Tyr         | Va<br>72 |
| Asp        | Thr        | Val        | Arg        | Ser<br>725 | Phe        | Arg        | Asp        | Arg        | Arg<br>730 | Tyr        | Glu              | Tyr        | Lys        | Thr<br>735  | Le       |
| Asn        | Lys        | Val        | Trp<br>740 | Lys        | Gly        | Lys        | Leu        | Ser<br>745 | Glu        | Ala        | Lys              | Ala        | Ser<br>750 | Gly         | As       |
| Leu        | He         | Lys<br>755 | He         | Gin        | Glu        | Ala        | His<br>760 | Asp        | Met        | Val        | Val              | Val<br>765 | Tyr        | Asp         | Se       |
| Leu        | GIn<br>770 | Leu        | Ala        | His        | Lys        | Cys<br>775 | lle        | Leu        | Asn        | Ser        | Phe<br>780       | Tyr        | Gly        | Tyr         | Va       |
| Met<br>785 | Arg        | Lys        | Gly        | Ala        | Arg<br>790 | Trp        | Tyr        | Ser        | Met        | Glu<br>795 | Met              | Ala        | Gly        | Val         | Va<br>80 |
| Thr        | Tyr        | Thr        | Gly        | Ala<br>805 | Lys        | He         | He         | Gln        | Asn<br>810 | Ala        | Arg              | Leu        | Leu        | l le<br>815 | Gl       |

| Arg         | He         | Gly         | Lys<br>820 | Pro                | Leu        | Glu        | Leu         | Asp<br>825  | Thr        | Asp        | Gly        | ile         | Trp<br>830 |            | Ala         |
|-------------|------------|-------------|------------|--------------------|------------|------------|-------------|-------------|------------|------------|------------|-------------|------------|------------|-------------|
| Leu         | Pro        | Gly<br>835  | Ser        | Phe                | Pro        | Glu        | Asn<br>840  | Phe         | Thr        | Phe        | Lys        | Thr<br>845  | He         | Asp        | Met         |
| Lys         | Lys<br>850 | Phe         | Thr        | lle                | Ser        | Tyr<br>855 | Pro         | Cys         | Val        | He         | Leu<br>860 | Asn         | Val        | Asp        | Val         |
| Ala<br>865  | Lys        | Asn         | Asn        | Ser                | Asn<br>870 | Asp        | GIn         | Tyr         | Gln        | Thr<br>875 | Leu        | Val         | Asp        | Pro        | Va I<br>880 |
| Arg         | Lys        | Thr         | Tyr        | <b>A</b> sn<br>885 | Ser        | Arg        | Ser         | Glu         | Cys<br>890 | Ser        | He         | Glu         | Phe        | Glu<br>895 | Val         |
| Asp         | Gly        | Pro         | Tyr<br>900 | Lys                | Ala        | Met        | lle         | l le<br>905 | Pro        | Ala        | Ser        | Lys         | Glu<br>910 | Glu        | Gly         |
| He          | Leu        | l le<br>915 | Lys        | Lys                | Arg        | Tyr        | Ala<br>920  | Val         | Phe        | Asn        | His        | Asp<br>925  | Gly        | Thr        | lle         |
| Ala         | Glu<br>930 | Leu         | Lys        | Gly                | Phe        | Glu<br>935 | Met         | Lys         | Arg        | Arg        | Gly<br>940 | Glu         | Leu        | Lys        | Leu         |
| l le<br>945 | Lys        | Val         | Phe        | Gln                | Ala<br>950 | Glu        | Leu         | Phe         | Asp        | Lys<br>955 | Phe        | Leu         | His        | Gly        | Ser<br>960  |
| Thr         | Leu        | Glu         | Glu        | Cys<br>965         | Tyr        | Ser        | Ala         | Val         | Ala<br>970 | Ala        | Val        | Ala         | Asn        | Arg<br>975 | Trp         |
| Leu         | Asp        | Leu         | Leu<br>980 | Glu                | Gly        | Gln        | Gly         | Lys<br>985  | Asp        | lle        | Ala        | Asp         | Ser<br>990 | Glu        | Leu         |
| Leu         | Asp        | Tyr<br>995  | lle        | Ser                | Glu        |            | Ser<br>1000 | Thr         | Met        | Ser        |            | Ser<br>1005 | Leu        | Ala        | Asp         |

| Tyr Gly<br>1010 |               | Gln         | Lys         |             | Cys<br>1015 | Ala                 | Vai                 | Thr          |             | Ala<br>1020 | Lys                  | Arg         | Leu         | Ala         |
|-----------------|---------------|-------------|-------------|-------------|-------------|---------------------|---------------------|--------------|-------------|-------------|----------------------|-------------|-------------|-------------|
| Asp Phe<br>1025 | e Leu         | Gly         |             | Thr<br>030  | Met         | Vai                 | Lys                 |              | Lys<br>1035 | Gly         | Leu                  | Arg         |             | GIn<br>1040 |
| Tyr lle         | val           |             | Arg<br>1045 | Glu         | Pro         | Glu                 |                     | Thr<br>1050  | Pro         | Val         | Ser                  |             | Arg<br>1055 | Ala         |
| Val Pro         |               | Ala<br>1060 | He          | Phe         | Gin         |                     | Asp<br>1065         | Asp          | Pro         | Glu         |                      | Lys<br>1070 | Phe         | Tyr         |
| Leu Gir         | Lys<br>1075   |             | Cys         | Lys         |             | Ser<br>1080         | Ser                 | Tyr          | Thr         |             | lle<br>1085          | Arg         | Ser         | He          |
| ile Asp<br>1090 |               | Met         | Tyr         |             | Lys<br>1095 | Gln                 | Arg                 | Leu          |             | Ser<br>1100 | Ala                  | lle         | Gln         | Lys         |
| Val IIe<br>1105 | e Thr         | lle         |             | Ala<br>110  | Ala         | Met                 | Gin                 |              | Va1<br>1115 | Ala         | Asn                  | Pro         |             | Leu<br>1120 |
| Arg Vai         | <b>A</b> rg   |             | Pro<br>1125 | Tyr         | Trp         | Leu                 |                     | Lys<br>  130 | Lys         | Val         | Cys                  |             | Lys<br>1135 | Phe         |
| Arg Gli         |               | Lys<br>1140 | lle         | Val         | Asp         |                     | Phe<br>11 <b>45</b> | Ser          | Ser         | Ala         |                      | Lys<br>1150 | Asp         | His         |
| Ser Thi         | - Thr<br>1155 |             | Asp         | Asn         |             | <b>V</b> al<br>1160 | Ala                 | Asp          | ile         |             | Glu<br>1 <b>16</b> 5 | Phe         | Cys         | Lys         |
| Glu Ası<br>1170 |               | Pro         | Ser         |             | Lys<br>1175 | Gly                 | Pro                 | Lys          |             | Val<br>1180 | Ala                  | Arg         | Ser         | Phe         |
| Glu Va<br>1185  | l Asp         | Arg         |             | His<br>1190 | Ser         | Glu                 | Gly                 |              | Gln<br>1195 | Gln         | Glu                  | Ser         |             | Asp<br>1200 |
| Pro Gli         | ı Phe         | Hie         | Aen         | ماا         | Sor         | len                 | Gln                 | Aen          | Vel         | Aen         | Lve                  | <b>A</b> en | Val         | Aen         |

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|                     | 1205                |                     | 1210                        | 1                   | 1215            |
|---------------------|---------------------|---------------------|-----------------------------|---------------------|-----------------|
|                     | Trp Leu Glu<br>1220 |                     | Lys Arg Lys<br>225          | Trp Lys Met<br>1230 | Thr Leu         |
| Thr Asn Lys<br>1235 | Lys Lys <b>A</b> rg | g Arg Phe 1<br>1240 | Asp Asp Leu                 | Lys Pro Cys<br>1245 | Asn Gin         |
| lie Asp Ala<br>1250 | His Lys Ile         | Asn Lys i<br>1255   |                             | Lys Gly Arg<br>1260 | Val Gly         |
| Val Gly Ser<br>1265 | Tyr Phe Arg         |                     | Glu Glu Ala<br>1275         | Leu Thr Ser         | Ser Tyr<br>1280 |
| Leu Gin Ile         | lle Gin Lei<br>1285 | ı Val Gin S         | Ser Pro Gln<br>1290         | Ser Gly Gln         | Phe Phe<br>1295 |
|                     | Val Val Gli<br>1300 |                     | Met Leu Lys<br>305          | lle Pro Leu<br>1310 | Thr lle         |
| Pro Arg Val<br>1315 | Phe Tyr II          | e Asn Ser  <br>1320 | Lys Ala Ser                 | lle Ala Gly<br>1325 | Asn Phe         |
| Thr Gly Lys<br>1330 | Cys lle As          | n Lys Ile  <br>1335 |                             | Gly Lys Pro<br>1340 | Cys Tyr         |
| Asn Leu Met<br>1345 | Glu Ala Ar          |                     | His <b>A</b> sn Thr<br>1355 | His Ile Leu         | Leu Leu<br>1360 |
| Val Asn ile         | Gin Giu As<br>1365  | o Gin Phe           | lle Lys Glu<br>1370         | Ser Lys Lys         | Leu Ala<br>1375 |
|                     | Ala Asp Pr<br>1380  |                     | Glu Gly ile<br>385          | Tyr Glu Thr<br>1390 |                 |
| Pro Leu Glu<br>1395 |                     | a lle Cys<br>1400   | Gin lie Giy                 | Cys Val Cys<br>1405 | Lys lle         |

| Glu Asp Thr<br>1410 |                       | Arg Asn Thi<br>1415 | r Gin Asp Giy<br>1420 | Trp Lys Leu Gly         |
|---------------------|-----------------------|---------------------|-----------------------|-------------------------|
| Glu Leu His         | s Arg lle Thr<br>1430 | Thr Thr Glu         | ı Cys Arg Tyr<br>1435 | Leu Glu Asn Ser<br>1440 |
| lle Pro Le          | u Val Tyr Leu<br>1445 | Tyr His Sen         | Thr Ser Thr<br>1450   | Gly Arg Ala Val<br>1455 |
| Tyr Val Leu         | ı Tyr Cys His<br>1460 | Ala Ser Lys         |                       | Val Val Val Val<br>1470 |
| Asn Pro Tyr<br>1475 |                       | Glu Leu Leu<br>1480 |                       | Leu Glu Arg Gln<br>1485 |
| Phe Arg Asp<br>1490 |                       | Glu Leu Ser<br>1495 | Pro Glu Pro<br>1500   | Phe Ser Trp Asp         |
| Gly lle Leu<br>1505 | Phe Gin Val           | Glu Tyr Va          | Asp His Pro<br>1515   | Glu Ala Ala Thr<br>1520 |
| Lys Phe Leu         | ı Gin Lys Ala<br>1525 | Leu Cys Glu         | ı Tyr ArgʻGlu<br>1530 | Glu Asn Cys Gly<br>1535 |
| Ala Thr Val         | Ala Val IIe<br>1540   | Glu Cys Pro         | •                     | Thr Thr Lys Glu<br>1550 |
| Gly Val Lys<br>1555 |                       | Asp Phe Pro<br>1560 | •                     | lle Pro Phe Asn<br>1565 |
| Asp Asp Asp<br>1570 |                       | Gin Pro Val<br>1575 | Ser Trp Gin<br>1580   | Arg Pro Ala Ala         |
| Lys lle Ala<br>1585 | ı Val Leu Arg<br>1590 | Cys Ala Ser         | Ala lle Gin<br>1595   | Trp Leu Asp Arg<br>1600 |

| Arg lie Ala Gi | n Ser Arg Tyr          | Ala His Val Pro              | Leu Gly Asn Phe Gly               |
|----------------|------------------------|------------------------------|-----------------------------------|
|                | 1605                   | 1610                         | 1615                              |
| Arg Asp Trp Le |                        | Val Asp IIe Phe<br>1625      | Leu Ser Arg Ala Leu<br>1630       |
| Arg Asp Gin Gi |                        | Trp Val Ser Asp              | Asn Gly Val Pro Asp               |
| 1635           |                        | 640                          | 1645                              |
| Leu Gly Asp II | le Asn Asn Glu<br>1655 | Glu Thr Phe Leu              | ı Ala Asp Glu Leu Gin<br>1660     |
| Glu Thr Ser Le | eu Leu Phe Pro<br>1670 | Gly Ala Tyr Arg<br>1675<br>, | Lys Val Ser Val Glu<br>1680       |
| Leu Lys Val Hi | is. Arg Leu Ala        | Val Asn Ala Leu              | ı Leu Lys Ser Asp Leu             |
|                | 1685                   | 1690                         | 1695                              |
| Val Ser Glu Mo |                        | Gly Phe Leu Gly              | Val Asn Ser Arg Gly               |
| 170            |                        | 1705                         | 1710                              |
| Ser Ser Leu As |                        | Ser Phe Asp Glo              | ı Asn Asn Gly Cys Ala             |
| 1715           |                        | 1720                         | 1725                              |
| Gin Ala Phe Al | rg Val Leu Lys<br>1735 | Gin Leu ile Lys              | s Arg Leu Leu His Asp<br>1740     |
| Ala Cys Asn S  | er Gly Asn lle<br>1750 | Tyr Ala Asp Sei              | r ile Leu Gin His Leu<br>5 _ 1760 |
| Ser Trp Trp L  | eu Arg Ser Pro         | Ser Ser Lys Lei              | u His Asp Pro Ala Leu             |
|                | 1765                   | 1770                         | 1775                              |
| His Leu Met L  |                        | Met Gin Lys Va<br>1785       | l Phe Ala Leu Leu Leu<br>1790     |

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| Thr         | -           | Leu<br>1795 | Arg         | Arg         | Leu         |             | Ala<br>1800 | He          | lle         | lle         |              | Ala<br>1805 | Asp         | Phe         | Ser         |
|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|--------------|-------------|-------------|-------------|-------------|
| -           | Val<br>1810 | lle         | He          | Asp         |             | Gly<br>1815 | Lys         | Phe         | Asp         |             | Ser<br>1820  | Ala         | Ala         | Lys         | Thr         |
| Tyr<br>1825 |             | Glu         | Ser         |             | Leu<br>1830 | Thr         | Val         | Met         |             | Ser<br>1835 | Arg          | Asp         | He          |             | Lys<br>1840 |
| Leu         | lle         | Leu         |             | Glu<br>1845 | Pro         | Val         | His         | _           | Trp<br>1850 | His         | Ser          | Leu         |             | Phe<br>1855 | Met         |
| Asp         | GIn         |             | Asn<br>1860 | Tyr         | Ala         | Gly         |             | Arg<br>1865 | Ala         | Thr         | Gly          | Asp         | Glu<br>1870 | lle         | Ser         |
| Gly         |             | Glu<br>1875 | Val         | Thr         | lle         |             | Pro<br>1880 | Lys         | Trp         | Ser         |              | Ala<br>1885 | Arg         | His         | Leu         |
|             | Glu<br>1890 | Tyr         | He          | Gln         |             | Asp<br>1895 | Phe         | lle         | He          |             | Va I<br>1900 | Ala         | Thr         | Phe         | He          |
| Phe<br>1905 |             | Pro         | Trp         |             | Phe<br>1910 | Ala         | Leu         | Glu         |             | Lys<br>1915 | Arg          | Gly         | Ser         |             | Glu<br>1920 |
| Ser         | Leu         | Glu         |             | Glu<br>1925 | Met         | Val         | Glu         |             | Leu<br>1930 | Lys         | Glu          | Gln         |             | Gly<br>1935 | Thr         |
| Arg         | Phe         |             | Ser<br>1940 | Met         | lle         | Val         |             | Lys<br>1945 | He          | Gly         | Asn          | lle         | Arg<br>1950 | Ser         | His         |
| lle         |             | Asp<br>1955 | lle         | Asn         | Val         |             | Asp<br>1960 | Ala         | Ser         | Trp         |              | Ser<br>1965 | Gly         | Gin         | Ala         |
|             | Lys<br>1970 | Gly         | Asp         | Tyr         |             | Phe<br>1975 | Glu         | Phe         | He          |             | ile<br>1980  | He          | Thr         | Ala         | Val         |
| Leu         | Ala         | Leu         | Asp         | Gln         | Asn         | Val         | Gin         | Gin         | Asp         | Val         | Leu          | Val         | Met         | Arg         | Lys         |

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<212> DNA

<213> Oryza sativa

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lle Leu Leu Lys Tyr lle Lys Val Lys Glu Cys Ala Ala Glu Ala Glu Phe IIe Asp Pro Gly Pro Ser Phe IIe Leu Pro Asn Val Ala Cys Ser Asn Cys Gly Ala Tyr Arg Asp Leu Asp Phe Cys Arg Asp Ser Ala Leu Leu Thr Glu Lys Glu Trp Ser Cys Ala Asp Pro Gln Cys Val Lys Ile Tyr Asp Lys Glu Gin lie Glu Ser Ser lie lie Gin Met Vai Arg Gin Arg Glu Arg Met Tyr Gln Leu Gln Asp Leu Vai Cys Asn Arg Cys Asn Gln Val Lys Ala Ala His Leu Thr Glu Gln Cys Glu Cys Ser Gly Ser Phe Arg Cys Lys Glu Ser Gly Ser Asp Phe His Lys Arg Ile Glu Ile Phe Leu Asp lle Ala Lys Arg Gin Lys Phe Arg Leu Leu Giu Giu Cys lle Ser Trp lle Leu Phe Ala Thr Ser Cys <210> 47 <211> 3706

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#### 64/236

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<210> 48 <211> 1105 <212> PRT <213> Oryza sativa

**<400> 48** 

Met Ser Ser Gly Gly Arg Gly Gly Lys Arg Arg Gly Ala Pro Pro 1 5 10 15

| Gly        | Pro        | Ser        | Gly<br>20  | Ala        | Ala        | Ala        | Lys         | Arg<br>25  | Ala        | His        | Pro        | Gly        | Gly<br>30  | Thr                | Pro        |
|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|--------------------|------------|
| Gln        | Pro        | Pro<br>35  | Pro        | Pro        | Ala        | Ala        | Thr<br>40   | Ala        | Ala        | Ala        | Pro        | Va I<br>45 | Ala        | Glu                | Glu        |
| Glu        | Asp<br>50  | Met        | Met        | Asp        | Glu        | Asp<br>55  | Val         | Phe        | Leu        | Asp        | Glu<br>60  | Thr        | lle        | Leu                | Ala        |
| Glu<br>65  | Asp        | Glu        | Glu        | Ala        | Leu<br>70  | Leu        | Leu         | Leu        | Asp        | Arg<br>75  | Asp        | Glu        | Ala        | Leu                | Ala<br>80  |
| Ser        | Arg        | Leu        | Ser        | Arg<br>85  | Trp        | Arg        | Arg         | Pro        | Ala<br>90  | Leu        | Pro        | Ala        | Asp        | Leu<br>95          | Ala        |
| Ser        | Gly        | Cys        | Ser<br>100 | Arg        | Asn        | Val        | Ala         | Phe<br>105 | Gin        | GIn        | Leu        | Glu        | lle<br>110 | Asp                | Tyr        |
| Val        | lle        | Gly<br>115 | Glu        | Ser        | His        | Lys        | Va I<br>120 | Leu        | Leu        | Pro        | Asn        | Ser<br>125 | Ser        | Gly                | Pro        |
| Ala        | Ala<br>130 | lle        | Leu        | Arg        | He         | Phe<br>135 | Gly         | Val        | Thr        | Arg        | Glu<br>140 | Gly        | His        | Ser                | .Val       |
| Cys<br>145 | Cys        | Glņ        | Val        | His        | Gly<br>150 | Phe        | Glu         | Pro        | Tyr        | Phe<br>155 | Tyr        | He         | Ser        | Cys                | Pro<br>160 |
| Met        | Gly        | Met        | Gly        | Pro<br>165 | Asp        | Asp        | lle         | Ser        | Arg<br>170 | Phe        | His        | Gln        | Thr        | Leu<br>1 <b>75</b> | Glu        |
| Gly        | Arg        | Met        | Lys<br>180 | Asp        | Ser        | Asn        | Arg         | Asn<br>185 | Ser        | Asn        | Val        | Pro        | Arg<br>190 | Phe                | Val        |
| Lys        | Arg        | lie<br>195 | Glu        | Leu        | Val        | Gin        | Lys<br>200  | Gin        | Thr        | lle        | Met        | His<br>205 | Tyr        | Gln                | Pro        |

| GIn        | GIn<br>210  | Ser         | Gln        | Pro        | Phe        | Leu<br>215 | Lys        | He          | Vai        | Val        | Ala<br>220 | Leu        | Pro        | Thr        | Met        |
|------------|-------------|-------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|
| Val<br>225 | Ala         | Ser         | Cys        | Arg        | Gly<br>230 | He         | Leu        | Glu         | Arg        | Gly<br>235 | l le       | Thr        | He         | Glu        | Gly<br>240 |
| Leu        | Gly         | Ser         | Lys        | Ser<br>245 | Phe        | Leu        | Thr        | Tyr         | Glu<br>250 | Ser        | Asn        | lle        | Leu        | Phe<br>255 | Ala        |
| Leu        | Arg         | Phe         | Met<br>260 | He         | Asp        | Cys        | Asn        | l le<br>265 | Val        | Gly        | Gly        | Asn        | Trp<br>270 | lle        | Glu        |
| Val        | Pro         | Ala<br>275  | Gly        | Lys        | Tyr        | Met        | Lys<br>280 | Ala         | Ala        | Arg        | lle        | Met<br>285 | Ser        | Tyr        | Cys        |
| Gin        | Leu<br>290  | Glu         | Leu        | Asp        | Cys        | Leu<br>295 | Tyr        | Ser         | Asp        | Leu        | Val<br>300 | Ser        | His        | Ala        | Ala        |
| Glu<br>305 | Gly         | Glu         | His        | Ser        | Lys<br>310 | Met        | Ala        | Pro         | Phe        | Arg<br>315 | lle        | Leu        | Ser        | Phe        | Asp<br>320 |
| lle        | Glu         | Cys         | Ala        | Gly<br>325 | Arg        | Lys        | Gly        | His         | Phe<br>330 | Pro        | Glu        | Pro        | Thr        | His<br>335 | Asp        |
| Pro        | Val         | He          | Gin<br>340 | lle        | Ala        | Asn        | Leu        | Va I<br>345 | Thr        | Leu        | Gln        | Gly        | Glu<br>350 | Gly        | Gln        |
| Pro        | Phe         | Va i<br>355 | Arg        | Asn        | Val        | Met        | Thr<br>360 | Leu         | Lys        | Ser        | Cys        | Ser<br>365 | Pro        | He         | Val        |
| Gly        | Va I<br>370 | Asp         | Val        | Met        | Ser        | Phe<br>375 | Asp        | Thr         | Glu        | Arg        | Asp<br>380 | Val        | Leu        | Leu        | Ala        |
| Trp<br>385 | Arg         | Asp         | Phe        | lle        | Arg<br>390 | Glu        | Val        | Asp         | Pro        | Asp<br>395 | lle        | lle        | He         | Gly        | Tyr<br>400 |
| Asn        | He          | Cys         | Lys        | Phe        | Asp        | Leu        | Pro        | Tyr         | Leu        | He         | Glu        | Arg        | Ala        | Glu        | Val        |

|             |            |             |             | 405        |            |             |            |            | 410        |            |            |            |            | 415        |            |
|-------------|------------|-------------|-------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| Leu         | Lys        | He          | Va I<br>420 | Glu        | Phe        | Pro         | lle        | Leu<br>425 | Giy        | Arg        | lle        | Arg        | Asn<br>430 | Ser        | Arg        |
| Val         | Arg        | Va I<br>435 | Arg         | Asp        | Thr        | Thr         | Phe<br>440 | Ser        | Ser        | Arg        | Gln        | Tyr<br>445 | Gly        | Met        | Arg        |
| Glu         | Ser<br>450 | Lys         | Asp         | Val        | Ala        | Va I<br>455 | Glu        | Gly        | Arg        | Val        | GIn<br>460 | Phe        | Asp        | Leu        | Leu        |
| GIn<br>465  | Ala        | Met         | Gln         | Arg        | Asp<br>470 | Tyr         | Lys        | Leu        | Ser        | Ser<br>475 | Tyr        | Ser        | Leu        | Asn        | Ser<br>480 |
| Val         | Ser        | Ala         | His         | Phe<br>485 | Leu        | Gly         | Glu        | Gin        | Lys<br>490 | Glu        | Asp        | Val        | His        | His<br>495 | Ser        |
| ile         | lle        | Ser         | Asp<br>500  | Leu        | Gln        | Asn         | Gly        | Asn<br>505 | Ser        | Glu        | Thr        | Arg        | Arg<br>510 | Arg        | Leu        |
| Ala         | Val        | Tyr<br>515  |             | Leu        | Lys        | Asp         | Ala<br>520 | Tyr        | Leu        | Pro        | Gln        | Arg<br>525 |            | Leu        | Asp        |
| Lys         | Leu<br>530 |             | Tyr         | lle        | Tyr        | Asn<br>535  | Tyr        | Val        | Glu        | Met        | Ala<br>540 | Arg        | Val        | Thr        | Gly        |
| Va i<br>545 |            | He          | Ser         | Phe        | Leu<br>550 | Leu         | Ser        | Arg        | Gly        | GIn<br>555 | Ser        | ile        | Lys        | Val        | Leu<br>560 |
| Ser         | Gln        | Leu         | Leu         | Arg<br>565 |            | Ala         | Lys        | Gln        | Lys<br>570 |            | Leu        | Val        | He         | Pro<br>575 |            |
| He          | Lys        | Gly         | Gin<br>580  |            | Ser        | Gly         | GIn        | Asp<br>585 |            | Phe        | Glu        | Gly        | Ala<br>590 | Thr        | Val        |
| Leu         | Glu        | Ala         | Arg         | Ala        | Gly        | Phe         | Tyr        | Glu        | Lys        | Pro        | lle        | Ala        | Thr        | Leu        | Asp        |

|            |            | 595        |            |            |            |            | 600        |            |            |            |                         | 605         |            |            |            |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------------------|-------------|------------|------------|------------|
|            | Ala<br>610 | Ser        | Leu        | Tyr        | Pro        | Ser<br>615 | lle        | Met        | Met        | Ala        | Tyr <sub>.</sub><br>620 | Asn         | Leu        | Cys        | Tyr        |
| Cys<br>625 | Thr        | Leu        | Val        | Pro        | Pro<br>630 | Glu        | Asp        | Ala        | Arg        | Lys<br>635 | Leu                     | Asn         | Leu        | Pro        | Pro<br>640 |
| Glu        | Ser        | Val        | Asn        | Lys<br>645 | Thr        | Pro        | Ser        | Gly        | Glu<br>650 | Thr        | Phe                     | Val         | Lys        | Pro<br>655 | Asp        |
| Val        | GIn        | Lys        | Gly<br>660 | He         | Leu        | Pro        | Glu        | lle<br>665 | Leu        | Glu        | Glu                     | Leu         | Leu<br>670 | Ala        | Ala        |
| Arg        | Lys        | Arg<br>675 | Ala        | Lys        | Ala        | Asp        | Leu<br>680 | Lys        | Glu        | Ala        | Lys                     | Asp<br>685  | Pro        | Phe        | Glu        |
| Arg        | Ala<br>690 | Val        | Leu        | Asp        | Gly        | Arg<br>695 | Gin        | Leu        | Ala        | Leu        | Lys<br>700              | lle         | Ser        | Ala        | Asn        |
| Ser<br>705 | Val        | Tyr        | Gly        | Phe        | Thr<br>710 | Gly        | Ala        | Thr        | Val        | Gly<br>715 | Gin                     | Leu         | Pro        | Cys        | Leu<br>720 |
| Glu        | lle        | Ser        | Ser        | Ser<br>725 | Val        | Thr        | Ser        | Tyr        | Gly<br>730 |            | GIn                     | Met         | He         | Glu<br>735 | His        |
| Thr        | Lys        | Lys        | Leu<br>740 |            | Glu        | Asp        | Lys        | Phe<br>745 | Thr        | Thr        | Leu                     | Gly         | Gly<br>750 |            | Glu        |
| His        | Asn        | Ala<br>755 |            | Val        | lle        | Tyr        | Gly<br>760 |            | Thr        | Asp        | Ser                     | Va I<br>765 |            | Val        | Gln        |
| Phe        | Gly<br>770 |            | Ser        | Thr        | Val        | Glu<br>775 |            | Ala        | Met        | Lys        | Leu<br>780              |             | Arg        | Glu        | Ala        |
| Ala<br>785 |            | Tyr        | lle        | Ser        | Gly<br>790 |            | Phe        | He         | Lys        | Pro<br>795 |                         | Lys         | Leu        | Glu        | Phe<br>800 |

| Glu        | Lys         | lle        | Tyr        | Phe<br>805 | Pro         | Tyr        | Leu        |            | 1 le<br>810 | Ser        | Lys        | Lys        | Arg        | Tyr<br>815 | Ala        |
|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|
| Gly        | Leu         | Tyr        | Trp<br>820 | Thr        | Asn         | Pro        | Glu        | Lys<br>825 | Phe         | Asp        | Lys        | Met        | Asp<br>830 | Thr        | Lys        |
| Gly        | lle         | Glu<br>835 | Thr        | Vai        | Arg         | Arg        | Asp<br>840 | Asn        | Cys         | Leu        | Leu        | Val<br>845 | Lys        | Asn        | Leu        |
| Val        | Thr<br>850  | Glu        | Cys        | Leu        | His         | Lys<br>855 | He         | Leu        | Val         | Asp        | Arg<br>860 | Asp        | Val        | Pro        | Gly        |
| Ala<br>865 | Val         | Gln        | Tyr        | Val        | Lys<br>870  | Asn        | Thr        | He         | Ser         | Asp<br>875 | Leu        | Leu        | Met        | Asn        | Arg<br>880 |
| Val        | Asp         | Leu        | Ser        | Leu<br>885 | Leu         | Val        | He         | Thr        | Lys<br>890  | Gly        | Leu        | Thr        | Lys        | Thr<br>895 | Gly        |
| Glu        | Asp         | Tyr        | Ala<br>900 | Val        | Lys         | Ala        | Ala        | His<br>905 | Val         | Glu        | Leu        | Ala        | Glu<br>910 | Arg        | Met        |
| Arg        | Lys         | Arg<br>915 | Asp        | Ala        | Ala         | Thr        | Ala<br>920 |            | Thr         | Val        | Gly        | Asp<br>925 |            | Val        | Pro        |
| Tyr        | Va I<br>930 |            | lle        | Lys        | Ala         | Ala<br>935 |            | Gly        | Ala         | Lys        | Ala<br>940 |            | Glu        | Arg        | Ser        |
| Glu<br>945 |             | Pro        | He         | Tyr        | Va I<br>950 |            | Asp        | Asn        | Asn         | 11e<br>955 |            | He         | Asp        | Pro        | GIn<br>960 |
| Tyr        | Tyr         | Leu        | Glu        | Asn<br>965 | Gln         | lle        | Ser        | Lys        | Pro<br>970  |            | Leu        | Arg        | lle        | Phe<br>975 |            |
| Pro        | lle         | Leu        | Lys<br>980 |            | Ala         | Ser        | Arg        | Glu<br>985 |             | Leu        | His        | Gly        | Ser<br>990 |            | Thr        |

Arg Ala Val Ser lie Ser Thr Pro Ser Asn Ser Gly lie Met Lys Phe 995 1000 1005

Ala Lys Lys Gln Leu Thr Cys Leu Gly Cys Lys Ala Val lie Ser Gly 1010 1015 1020

Ser Asn Gln Thr Leu Cys Phe His Cys Lys Gly Arg Glu Ala Glu Leu 1025 1030 1035 1040

Tyr Cys Lys Thr Val Gly Asn Val Ser Glu Leu Glu Met Leu Phe Gly 1045 1050 1055

Arg Leu Trp Thr Gln Cys Gln Glu Cys Gln Gly Ser Leu His Gln Asp 1060 1065 1070

Val Leu Cys Thr Ser Arg Asp Cys Pro I le Phe Tyr Arg Arg Lys 1075 1080 1085

Ala Gin Lys Asp Met Ala Giu Ala Arg Val Gin Leu Gin Arg Trp Asp 1090 1095 1100

Phe

1105

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<213> Glycine max

#### <400> 49

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| gaactcgtct | ggacctgtcg | ccattatcag | aatatttgga   | gttactaagg | aaggacacag   | 420  |
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| gggtccagat | gatatctccc | attttcatca | aactctcgag   | ggaaggatga | gagaagccaa   | 540  |
| tagaaacagt | aatgtgggaa | aattcgttcg | ccgtattgaa   | atggtgcaga | gaaggagtat   | 600  |
| tatgtactat | cagcaatcca | attcccaacc | ctttctcaaa   | attgtagttg | cactcccaac   | 660  |
| aatggttgcc | agctgccgtg | gtattcttga | taggggtatt   | caacttgatg | gtctgggaat   | 720  |
| gaagagcttc | ttgacttatg | aaagcaatgt | actttttgcc   | cttcgcttca | tgattgattg   | 780  |
| taacatagtt | ggtggaaatt | ggattgggat | tcctgccgga   | aaatataaga | aaacagcgaa   | 840  |
| aagcttgtct | tactgccagt | tagagtttga | ttgcttgtat   | totgaattga | ttagtcatgc   | 900  |
| tccagaaggg | gaatattcaa | agatggctcc | gtttcgcatt   | ttgagttttg | acatcgagtg   | 960  |
| tgctggtcgt | aaaggtcatt | ttcctgagcc | tacccatgat   | cctgttatcc | agattgctaa   | 1020 |
| tttggttact | ttacaaggag | aagaccagcc | atttattcgt   | aatgtgatga | cccttaaatc   | 1080 |
| atgttctcct | atcgttggtg | ttgatgtgat | gccatttgaa   | acagaaagag | aagtcctgct   | 1140 |
| ggcttggagg | gattttattc | gtgaagtgga | ccctgatatt   | attattggat | acaacatttg   | 1200 |
| caaatttgac | ttgccatatc | ttattgagag | agctttgaac   | ctgaagatag | cagaatttcc   | 1260 |
|            |            |            |              |            | tctcatcaag   |      |
| gcagtacgga | accagggaaa | gtaaagaagt | tgcagtagaa   | gggagagtta | cgtttgattt   | 1380 |
|            |            |            |              |            | ctgtgtcatc   |      |
|            |            |            |              |            | atcttcagaa   |      |
|            |            |            |              |            | catatetece   |      |
|            |            |            |              |            | ctcgagtaac   |      |
|            |            |            |              |            | tttctcaact   |      |
|            |            |            |              |            | ctgggtctga   |      |
|            |            |            |              |            | atgaaaaacc   |      |
| _          |            |            |              |            | ataacttatg   |      |
|            |            |            |              |            | cagagtctgt   |      |
| _          |            |            |              |            | gaatacttcc   |      |
|            |            |            |              |            | taaaggaggc   |      |
|            |            |            |              |            | agattagtgc   |      |
|            |            |            |              | •          | tagagatatc   |      |
|            |            |            |              |            | ttgtggaaga   |      |
|            |            |            |              |            | gagacacaga   |      |
|            |            |            |              |            | tggggagaga   |      |
|            |            |            |              |            | : ttgagaaggt |      |
|            |            |            |              |            | ggacaaaacc   |      |
|            |            |            |              |            | acaattgttt   |      |
| attggtcaaa | aacctggtga | acgattgcct | : tcacaaaata | ttgattgaca | gggacattco   | 2580 |

tggggcagtc cagtatgtca agaatgcaat ttcagatctt ctcatgaatc gtatggactt 2640 atcacttctg gttattacaa agggtttaac gaagacagga gatgattacg aagtaaaggc 2700 agctcatgtt gaacttgctg aaaggatgcg caagcgagat gctgccactg ctccaaatgt 2760 tggagacaga gtaccatatg ttattattaa agctgcaaaa ggtgcaaagg catatgagag 2820 atcagaggat cctatctatg tgctagagaa caacataccc atagatcctc attactatct 2880 tgagaatcaa attagcaagc caattctgag aatttttgag ccaattctga agaatgctag 2940 caaagagett eteeatggaa gteatacaag atetattet atttetacae egteaaacag 3000 tggcatattg agatttgcta agaaacagct acctgcattg gttgtaaagc tttacttggc 3060 aagggttatc acactetetg tteacattge aaaggaaggg aggetgaget gtactgtaaa 3120 acagtatoto aagtgtotga gotggagatg otttttggga ggttgtggac acagtgtoag 3180 gagtgccaag gttcacttca tcaggatgtt ctctgcacca gtcgggattg tccaattttc 3240 tatcgacgaa aaaaggcaca gaaagatatg ggtgaagcaa agttgcaatt ggacagatgg 3300 aacttotaag ttttgccaag aatttgacct tgcggatoto ttcgaaccaa tggacacaaa 3360 tacaatctgg tgtttgccac aatcctgaca tttgtaatgt gagtaaaagc ccacaatttg 3420 3427 tttactg

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<211> 1088

<212> PRT

<213> Glycine max

<400> 50

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Gin Arg Gin Ala Leu Ala Asn Arg Leu Ser Lys Trp Thr Arg Pro Pro 35 40 45

Leu Ser Ala Gly Tyr Val Ala Gln Ser Arg Ser Val Leu Phe Gln Gln
50 55 60

Leu Glu lle Asp Tyr Val lle Ala Glu Ser His Gly Glu Leu Leu Pro

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| 65          |             |            |            |                  | 70         |            |            |            |            | 75         |                  |            |            |            | 80         |
|-------------|-------------|------------|------------|------------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|
| <b>A</b> sn | Ser         | Ser        | Gly        | Pro<br>85        | Val        | Ala        | lle        | lle        | Arg<br>90  | He         | Phe <sub>.</sub> | Gly        | Val        | Thr<br>95  | Lys        |
| Glu         | Gly         | His        | Ser<br>100 | Val              | Cys        | Cys        | Asn        | Val<br>105 | His        | Gly        | Phe              | Glu        | Pro<br>110 | Tyr        | Phe        |
| Tyr         | lle         | Cys<br>115 | Cys        | Pro              | Pro        | Gly        | Met<br>120 | Gly        | Pro        | Asp        | Asp              | lle<br>125 | Ser        | His        | Phe        |
| His         | GIn<br>130  | Thr        | Leu        | Glu              | Gly        | Arg<br>135 | Met        | Arg        | Glu        | Ala        | Asn<br>140       | Arg        | Asn        | Ser        | Asn        |
| Va I<br>145 | Gly         | Lys        | Phe        | Val              | Arg<br>150 | Arg        | He         | Glu        | Met        | Val<br>155 | Gln              | Arg        | Arg        | Ser        | lle<br>160 |
| Met         | Tyr         | Tyr        | Gln        | Gin<br>165       | Ser        | Asn        | Ser        | Gln        | Pro<br>170 | Phe        | Leu              | Lys        | He         | Val<br>175 | Val        |
| Ala         | Leu         | Pro        | Thr<br>180 | Met <sub>.</sub> | Val        | Ala        | Ser        | Cys<br>185 | Arg        | Gly        | ile              | Leu        | Asp<br>190 | Arg        | Gly        |
| He          | Gin         | Leu<br>195 | Asp        | Gly              | Leu        | Gly        | Met<br>200 | Lys        | Ser        | Phe        | Leu              | Thr<br>205 | Tyr        | Glu        | Ser        |
| Asn         | Va I<br>210 | Leu        | Phe        | Ala              | Leu        | Arg<br>215 | Phe        | Met        | lle        | Asp        | Cys<br>220       | Asn        | lle        | Val        | Gly        |
| Gly<br>225  | Asn         | Trp        | lle        | Gly              | 11e<br>230 | Pro        | Ala        | Giy        | Lys        | Tyr<br>235 | •                | Lys        | Thr        | Ala        | Lys<br>240 |
| Ser         | Leu         | Ser        | Tyr        | Cys<br>245       |            | Leu        | Glu        | Phe        | Asp<br>250 |            | Leu              | Tyr        | Ser        | Glu<br>255 | Leu        |
| lle         | Ser         | His        | Ala<br>260 |                  | Glu        | Gly        | Glu        | Tyr<br>265 |            | Lys        | Met              | Ala        | Pro<br>270 | Phe        | Arg        |

| lle         | Leu         | Ser<br>275  | Phe        | Asp         | lle         | Glu         | Cys<br>280 | Ala        | Gly         | Arg        | Lys        | Gly<br>285 | His        | Phe        | Pro        |
|-------------|-------------|-------------|------------|-------------|-------------|-------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|
| Glu         | Pro<br>290  | Thr         | His        | Asp         | Pro         | Va I<br>295 | He         | Gln        | lle         | Ala        | Asn<br>300 | Leu        | Val        | Thr        | Leu        |
| GIn<br>305  | Gly         | Glu         | Asp        | Gin         | Pro<br>310  | Phe         | lle        | Arg        | Asn         | Val<br>315 | Met        | Thr        | Leu        | Lys        | Ser<br>320 |
| Cys         | Ser         | Pro         | He         | Va I<br>325 | Gly         | <b>V</b> al | Asp        | Val        | Met<br>330  | Pro        | Phe        | Glu        | Thr        | Glu<br>335 | Arg        |
| Glu         | Val         | Leu         | Leu<br>340 | Ala         | Trp         | Arg         | Asp        | Phe<br>345 | He          | Arg        | Glu        | Val        | Asp<br>350 | Pro        | Asp        |
| He          | lle         | l le<br>355 | Gly        | Tyr         | Asn         | lle         | Cys<br>360 | Lys        | Phe         | Asp        | Leu        | Pro<br>365 | Tyr        | Leu        | He         |
| Glu         | Arg<br>370  | Ala         | Leu        | Asn         | Leu         | Lys<br>375  | He         | Ala        | Glu         | Phe        | Pro<br>380 | lle        | Leu        | Gly        | Arg        |
| l le<br>385 |             | Asn         | Ser        | Arg         | Va I<br>390 | Arg         | Val        | Lys        | Asp         | Thr<br>395 | Thr        | Phe        | Ser        | Ser        | Arg<br>400 |
| Gin         | Tyr         | Gly         | Thr        | Arg<br>405  | Glu         | Ser         | Lys        | Glu        | Va I<br>410 | Ala        | Val        | Glu        | Gly        | Arg<br>415 | Val        |
| Thr         | Phe         | Asp         | Leu<br>420 |             | Gln         | Val         | Met        | GIn<br>425 |             | Asp        | Tyr        | Lys        | Leu<br>430 | Ser        | Ser        |
| Tyr         | Ser         | L.eu<br>435 |            | Ser         | Val         | Ser         | Ser<br>440 | His        | Phe         | Leu        | Ser        | Glu<br>445 |            | Lys        | Glu        |
| Asp         | Va I<br>450 |             | His        | Ser         | He          | 11e<br>455  |            | Asp        | Leu         | Gln        | Asn<br>460 |            | Asn        | Ala        | Glu        |

| Thr<br>465 | Arg        | Arg        | Arg        | Leu        | Ala<br>470 | Val        | Tyr         | Cys         | Leu        | Lys<br>475 | Asp        | Ala        | Tyr        | Leu        | Pro<br>480 |
|------------|------------|------------|------------|------------|------------|------------|-------------|-------------|------------|------------|------------|------------|------------|------------|------------|
| Gln        | Arg        | Leu        | Leu        | Asp<br>485 | Lys        | Leu        | Met         | Phe         | lle<br>490 | Tyr        | Asn        | Tyr        | Val        | Glu<br>495 | Met        |
| Ala        | Arg        | Val        | Thr<br>500 | Gly        | Val        | Pro        | He          | Ser<br>505  | Phe        | Leu        | Leu        | Ser        | Arg<br>510 | Gly        | Gln        |
| Ser        | He         | Lys<br>515 | Val        | Leu        | Ser        | Gln        | Leu<br>520  | Leu         | Arg        | Arg        | Ala        | Arg<br>525 | Gin        | Lys        | Asn        |
| Leu        | Val<br>530 | He         | Pro        | Asn        | Ala        | Lys<br>535 | Gln         | Ala         | Gly        | Ser        | Glu<br>540 | Gln        | Gly        | Thr        | Phe        |
| Glu<br>545 | Gly        | Ala        | Thr        | Val        | Leu<br>550 | Glu        | Ala         | Arg         | Ala        | Gly<br>555 | Phe        | Tyr        | Glu        | Lys        | Pro<br>560 |
| He         | Ala        | Thr        | Leu        | Asp<br>565 | Phe        | Ala        | Ser         | Leu         | Tyr<br>570 | Pro        | Ser        | lle        | Met        | Met<br>575 | Ala        |
| Tyr        | Asn        | Leu        | Cys<br>580 | Tyr        | Cys        | Thr        | Leu         | Va I<br>585 | lle        | Pro        | Glü        | Asp        | Ala<br>590 | Arg        | Lys        |
| Leu        | Asn        | lle<br>595 | Pro        | Pro        | Glu        | Ser        | Va I<br>600 | <b>As</b> n | Arg        | Thr        | Pro        | Ser<br>605 | Gly        | Glu        | Thr        |
| Phe        | Val<br>610 | Lys        | Ser        | Asn        | Leu        | GIn<br>615 | Lys         | Gly         | lle        | Leu        | Pro<br>620 | Glu        | lle        | Leu        | Glu        |
| Glu<br>625 | Leu        | Leu        | Thr        | Ala        | Arg<br>630 |            | Arg         | Ala         | Lys        | Ala<br>635 |            | Leu        | Lys        | Glu        | Ala<br>640 |
| Lys        | Asp        | Pro        | Leu        | Glu<br>645 | Lys        | Ala        | Val         | Leu         | Asp<br>650 |            | Arg        | Gln        | Leu        | Ala<br>655 | Leu        |

| Lys        | He         | Ser        | Ala<br>660 | Asn        | Ser        | Val        | Tyr         | Gly<br>665  | Phe        | Thr         | Gly        | Ala        | Thr<br>670 | lle          | Gly        |
|------------|------------|------------|------------|------------|------------|------------|-------------|-------------|------------|-------------|------------|------------|------------|--------------|------------|
| Gln        | Leu        | Pro<br>675 | Cys        | Leu        | Glu        | lie        | Ser<br>680  | Ser         | Ser        | Val         | Thr        | Ser<br>685 | Tyr        | Gly          | Arg        |
| GIn        | Met<br>690 | He         | Glu        | His        | Thr        | Lys<br>695 | Lys         | Leu         | Val        | Glu         | Asp<br>700 | Lys        | Phe        | Thr          | Thr        |
| Leu<br>705 | Asn        | Gly        | Tyr        | Glu        | His<br>710 | Asn        | Ala         | Glu         | Vai        | 1 le<br>715 | Tyr        | Gly        | Asp        | Thr          | Asp<br>720 |
| Ser        | Val        | Met        | Val        | GIn<br>725 | Phe        | Gly        | Val         | Ser         | Ala<br>730 | Val         | Glu        | Glu        | Ala        | Met<br>735   | Asn        |
| Leu        | Giy        | Arg        | Glu<br>740 | Ala        | Ala        | Glu        | His         | 1 le<br>745 | Ser        | Gly         | Thr        | Phe        | Thr<br>750 | Lys          | Pro        |
| He         | Lys        | Leu<br>755 | Glu        | Phe        | Glu        | Lys        | Va I<br>760 | Tyr         | Tyr        | Pro         | Tyr        | Leu<br>765 | Leu        | lle          | Ser        |
| Lys        | Lys<br>770 |            | Tyr        | Ala        | Gly        | Leu<br>775 | Phe         | Trp         | Thr        | Lys         | Pro<br>780 |            | Asn        | Phe          | Asp        |
| Lys<br>785 |            | Asp        | Thr        | Lys        | Gly<br>790 | lle        | Glu         | Thr         | Val        | Arg<br>795  |            | Asp        | Asn        | Cys          | Leu<br>800 |
| Leu        | Val        | Lys        | Asn        | Leu<br>805 |            | Asn        | Asp         | Cys         | Leu<br>810 |             | Lys        | He         | Leu        | 1 l e<br>815 | Asp        |
| Arg        | Asp        | lle        | Pro<br>820 |            | Ala        | Val        | GIn         | Tyr<br>825  |            | Lys         | Asn        | Ala        | lle<br>830 |              | Asp        |
| Leu        | Leu        | Met<br>835 |            | Arg        | Met        | Asp        | Leu<br>840  |             | Leu        | Leu         | Val        | lle<br>845 |            | Lys          | Gly        |
| Leu        | Thr        | Lys        | Thr        | Gly        | Asp        | Asp        | Tyr         | Glu         | Val        | Lys         | Ala        | Ala        | His        | Val          | Glu        |

|            | 850         |            |            |             |               | 855         |             |             |             |             | 860              |             |            |             |             |
|------------|-------------|------------|------------|-------------|---------------|-------------|-------------|-------------|-------------|-------------|------------------|-------------|------------|-------------|-------------|
| Leu<br>865 | Ala         | Glu        | Arg        | Met         | Arg<br>870    | Lys         | Arg         | Asp         | Ala         | Ala<br>875  | Thr <sub>.</sub> | Ala         | Pro        | Asn         | Val<br>880  |
| Gly        | Asp         | Arg        | Val        | Pro<br>885  | Tyr           | Val         | lle         | He          | Lys<br>890  | Ala         | Ala              | Lys         | Gly        | Ala<br>895  | Lys         |
| Ala        | Tyr         | Glu        | Arg<br>900 | Ser         | Glu           | Asp         | Pro         | l le<br>905 | Tyr         | Val         | Leu              | Glu         | Asn<br>910 | Asn         | He          |
| Pro        | ile         | Asp<br>915 | Pro        | His         | Tyr           | Tyr         | Leu<br>920  | Glu         | Asn         | Gln         | lle              | Ser<br>925  | Lys        | Pro         | lle         |
| Leu        | Arg<br>930  | lle        | Phe        | Glu         | Pro           | l le<br>935 | Leu         | Lys         | Asn         | Ala         | Ser<br>940       | Lys         | Glu        | Leu         | Leu         |
| His<br>945 |             | Ser        | His        | Thr         | Arg<br>950    |             | He          | Ser         | He          | Ser<br>955  | Thr              | Pro         | Ser        | Asn         | Ser<br>960  |
| Gly        | He          | Leu        | Arg        | Phe<br>965  |               | Lys         | Lys         | Gln         | Leu<br>970  |             | Ala              | Leu         | Val        | Va I<br>975 |             |
| Leu        | Tyr         | Leu        | Ala<br>980 |             | Val           | He          | Thr         | Leu<br>985  |             | Val         | His              | ile         | Ala<br>990 |             | Glu         |
| Gly        | Arg         | Leu<br>995 |            | Cys         | Thr           |             | Lys<br>1000 | Gin         | Tyr         | Leu         | Lys              | Cys<br>1005 |            | Ser         | Trp         |
| Arg        | Cys<br>1010 |            | Leu        | Gly         |               | Cys<br>1015 |             | His         | Ser         |             | Arg<br>1020      |             | Ala        | Lys         | Val         |
| His<br>102 |             | : He       | . Arg      | , Met       | : Phe<br>1030 |             | · Ala       | Pro         | Val         | Gly<br>1035 | lle<br>i         | ∤ Val       | Gin        | Phe         | Ser<br>1040 |
| Пe         | e Asp       | Glu        | ı Lys      | Arg<br>1045 |               | Arg         | , Lys       | He          | Trp<br>1050 |             | Lys              | Glr         | Ser        | Cys         |             |

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#### 79/236

Trp Thr Asp Gly Thr Ser Lys Phe Cys Gln Glu Phe Asp Leu Ala Asp 1060 1065 1070

Leu Phe Glu Pro Met Asp Thr Asn Thr lle Trp Cys Leu Pro Gln Ser 1075 1080 1085

<210> 51

<211> 3435

<212> DNA

⟨213⟩ Homo sapiens

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| tacccgtcca | tcatgatggc | ccacaacctg | tgttacacca | cactccttcg | gcccgggact | 1920 |
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| accagogoca | aggtggtgta | tggtgacact | gactccgtca | tgtgccgatt | cggcgtgtcc | 2340 |
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| Arg Ala Arg Gly Gly Leu Trp Asp Asp Asp Asp Ala Pro Arg Pro Ser<br>20 25 30    |      |
| Gin Phe Giu Giu Asp Leu Ala Leu Met Giu Giu Met Giu Ala Giu His<br>35 40 45    |      |
| Arg Leu Gin Giu Giu Giu Giu Giu Leu Gin Ser Val Leu Giu Giy 50 55 60           |      |
| Val Ala Asp Gly Gln Val Pro Pro Ser Ala IIe Asp Pro Arg Trp Leu<br>65 70 75 80 |      |
| Arg Pro Thr Pro Pro Ala Leu Asp Pro Gin Thr Giu Pro Leu I le Phe<br>85 90 95   |      |
| Gin Gin Leu Giu ile Asp His Tyr Val Gly Pro Ala Gin Pro Val Pro<br>100 105 110 |      |
| Gly Gly Pro Pro Pro Ser His Gly Ser Val Pro Val Leu Arg Ala Phe 115 120 125    |      |
| Gly Val Thr Asp Glu Gly Phe Ser Val Cys Cys His IIe His Gly Phe<br>130 135 140 |      |
| Ala Pro Tyr Phe Tyr Thr Pro Ala Pro Pro Gly Phe Gly Pro Glu His                |      |

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| 145        |             |            |             |            | 150        |             |            |             |            | 155        |                   |            |            |            | 160        |
|------------|-------------|------------|-------------|------------|------------|-------------|------------|-------------|------------|------------|-------------------|------------|------------|------------|------------|
| Met        | Gly         | Asp        | Leu         | Gln<br>165 | Arg        | Glu         | Leu        | <b>A</b> sn | Leu<br>170 | Ala        | l le <sub>.</sub> | Asn        | Arg        | Asp<br>175 | Ser        |
| Arg        | Gly         | Gly        | Arg<br>180  | Glu        | Leu        | Thr         | Gly        | Pro<br>185  | Ala        | Val        | Leu               | Ala        | Val<br>190 | Glu        | Leu        |
| Cys        | Ser         | Arg<br>195 | Glu         | Ser        | Met        | Phe         | Gly<br>200 | Tyr         | His        | Gly        | His               | Gly<br>205 | Pro        | Ser        | Pro        |
| Phe        | Leu<br>210  | Arg        | He          | Thr        | Val        | Ala<br>215  | Leu        | Pro         | Arg        | Leu        | Va I<br>220       | Ala        | Pro        | Ala        | Arg        |
| Arg<br>225 | Leu         | Leu        | Glu         | Gln        | Gly<br>230 | lle         | Arg        | Val         | Ala        | Gly<br>235 | Leu               | Gly        | Thr        | Pro        | Ser<br>240 |
| Phe        | Ala         | Pro        | Tyr         | Glu<br>245 | Ala        | Asn         | Val        | Asp         | Phe<br>250 |            | lle               | Arg        | Phe        | Met<br>255 | Val        |
| Asp        | Thr         | Asp        | l le<br>260 | Val        | Gly        | Cys         | Asn        | Trp<br>265  | Leu        | Glu        | Leu               | Pro        | Ala<br>270 | Gly        | Lys        |
| Tyr        | Ala         | Leu<br>275 | Arg         | Leu        | Lys        | Glu         | Lys<br>280 | Ala         | Thr        | GIn        | Cys               | GIn<br>285 | Leu        | Glu        | Ala        |
| Asp        | Va I<br>290 | Leu        | Trp         | Ser        | Asp        | Va I<br>295 |            | Ser         | His        | Pro        | Pro<br>300        | Glu        | Gly        | Pro        | Trp        |
| GIn<br>305 |             | lle        | Ala         | Pro        | Leu<br>310 |             | Val        | Leu         | Ser        | Phe<br>315 | •                 | He         | Glu        | Cys        | Ala<br>320 |
| Gly        | Arg         | Lys        | Gly         | lle<br>325 |            | Pro         | Glu        | Pro         | Glu<br>330 |            | Asp               | Pro        | Val        | 11e<br>335 | GIn        |
| Пe         | Cys         | Ser        | Leu<br>340  |            | Leu        | Arg         | Trp        | Gly<br>345  |            | Pro        | Glu               | Pro        | Phe<br>350 | Leu        | Arg        |

| _eu         | Ala         | Leu<br>355 | Thr        | Leu        | Arg        | Pro         | Cys<br>360 | Ala        | Pro        | lle        |            | Gly<br>365 | Ala        | Lys         | Val        |
|-------------|-------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|
| 3In         | Ser<br>370  | Tyr        | Glu        | Lys        | Glu        | Glu<br>375  | Asp        | Leu        | Leu        | GIn        | Ala<br>380 | Тгр        | Ser        | Thr         | Phe        |
| l le<br>385 | Arg         | He         | Met        | Asp        | Pro<br>390 | Asp         | Val        | lle        | Thr        | Gly<br>395 | Tyr        | Asn        | He         | GIn         | Asn<br>400 |
| Phe         | Asp         | Leu        | Pro        | Tyr<br>405 | Leu        | lle         | Ser        | Arg        | Ala<br>410 | Gln        | Thr        | Leu        | Lys        | Va I<br>415 | Gln        |
| Thr         | Phe         | Pro        | Phe<br>420 | Leu        | Gly        | Arg         | Val        | Ala<br>425 | Gly        | Leu        | Cyś        | Ser        | Asn<br>430 | lle         | Arg        |
| Asp         | Ser         | Ser<br>435 | Phe        | Gin        | Ser        | Lys         | GIn<br>440 | Thr        | Gly        | Arg        | Arg        | Asp<br>445 | Thr        | Lys         | Val        |
| Val         | Ser<br>450  | Met        | Vai        | Gly        | Arg        | Va I<br>455 | Gln        | Met        | Asp        | Met        | Leu<br>460 | Gin        | Val        | Leu         | Leu        |
| Arg<br>465  | Glu         | Tyr        | Lys        | Leu        | Arg<br>470 | Ser         | Tyr        | Thr        | Leu        | Asn<br>475 | Ala        | Val        | Ser        | Phe         | His<br>480 |
| Phe         | Leu         | Gly        | Glu        | GIn<br>485 | Lys        | Glu         | Asp        | Val        | GIn<br>490 |            | Ser        | He         | lle        | Thr<br>495  | Asp        |
| Leu         | Gln         | Asn        | Gly<br>500 | Asn        | Asp        | Gln         | Thr        | Arg<br>505 |            | Arg        | Leu        | Ala        | Val<br>510 |             | Cys        |
| Leu         | Lys         | Asp<br>515 |            | Tyr        | Leu        | Pro         | Leu<br>520 |            | Leu        | Leu        | Glu        | Arg<br>525 |            | Met         | Val        |
| Leu         | Va I<br>530 |            | Ala        | Val        | Glu        | Met<br>535  |            | Arg        | Val        | Thr        | Gly<br>540 |            | Pro        | Leu         | Ser        |

| Tyr<br>545 | Leu         | Leu        | Ser        | Arg        | Gly<br>550 | GIn         | GIn        | Val        | Lys        | Va I<br>555 | Vai         | Ser         | Gln        | Leu         | <b>560</b>  |
|------------|-------------|------------|------------|------------|------------|-------------|------------|------------|------------|-------------|-------------|-------------|------------|-------------|-------------|
| Arg        | GIn         | Ala        | Met        | His<br>565 | Glu        | Gly         | Leu        | Leu        | Met<br>570 | Pro         | Val         | Val         | L.ys       | Ser<br>575  | Glu         |
| âly        | Gly         | ,<br>Glu   | Asp<br>580 | Tyr        | Thr        | Gly         | Ala        | Thr<br>585 | Val        | He          | Glu         | Pro         | Leu<br>590 | Lys         | Gly         |
| Tyr        | Tyr         | Asp<br>595 | Val        | Pro        | lle        | Ala         | Thr<br>600 | Leu        | Asp        | Phe         | Ser         | Ser<br>605  | Leu        | Tyr         | Pro         |
| Ser        | l le<br>610 | Met        | Met        | Ala        | His        | Asn<br>615  | Leu        | Cys        | Tyr        | Thr         | Thr<br>620  | Leu         | Leu        | Arg         | Pro         |
| Gly<br>625 | Thr         | Ala        | Gln        | Lys        | Leu<br>630 | Gly         | Leu        | Thr        | Glu        | Asp<br>635  | Gln         | Phe         | lle        | Arg         | Thr<br>640  |
| Pro        | Thr         | Gly        | Asp        | Glu<br>645 | Phe        | Val         | Lys        | Thr        | Ser<br>650 |             | Arg         | Lys         | Gly        | Leu<br>655  | Leu         |
| Pro        | Gin         | lle        | Leu<br>660 |            | Asn        | Leu         | Leu        | Ser<br>665 | Ala        | Arg         | Lys         | Arg         | Ala<br>670 | Lys         | Ala         |
| Glu<br>•   | Leu         | Ala<br>675 | Lys        | Glu        | Thr        | Asp         | Pro<br>680 |            | Arg        | Arg         | Gln         | Va I<br>685 |            | Asp         | Gly         |
| Arg        | GIn<br>690  |            | Ala        | Leu        | Lys        | Va I<br>695 |            | Ala        | Asn        | Ser         | Va !<br>700 | Tyr         | Gly        | Phe         | Thr         |
| Gly<br>705 |             | Gln        | Val        | Gly        | Lys<br>710 |             | Pro        | Cys        | Leu        | Glu<br>715  |             | Ser         | GIn        | Ser         | Va I<br>720 |
| Thr        | Gly         | Phe        | Gly        | 725        |            | Met         | lle        | Glu        | Lys<br>730 |             | Lys         | GIn         | Leu        | Va I<br>735 | Glu         |
| Ser        | Lys         | Tyr        | Thr        | Val        | Glu        | Asn         | Gly        | Tyr        | Ser        | Thr         | Ser         | Ala         | Lys        | Val         | Val         |

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|            |            |            | 740        |            |            |            |             | 745        |            |            |            |             | 750        |            |            |
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| Tyr        | Gly        | Asp<br>755 | Thr        | Asp        | Ser        | Val        | Met<br>760  | Cys        | Arg        | Phe        | Gly        | Va I<br>765 | Ser        | Ser        | Val        |
| Ala        | Glu<br>770 | Ala        | Met        | Ala        | Leu        | Gly<br>775 | Gly         | Glu        | Ala        | Ala        | Asp<br>780 | Trp         | Val        | Ser        | Gly        |
| His<br>785 | Phe        | Pro        | Ser        | Pro '      | 11e<br>790 | Arg        | Leu         | Glu        | Phe        | Glu<br>795 | Lys        | Val         | Tyr        | Phe        | Pro<br>800 |
| Tyr        | Leu        | Leu        | He         | Ser<br>805 | Lys        | Lys        | Arg         | Tyr        | Ala<br>810 |            | Leu        | Leu         | Phe        | Ser<br>815 | Ser        |
| Arg        | Pro        | Asp        | Ala<br>820 | His        | Asp        | Arg        | Met         | Asp<br>825 | Cys        | Lys        | Gly        | Leu         | Glu<br>830 | Ala        | Val        |
| Arg        | Arg        | Asp<br>835 | Asn        | Cys        | Pro        | Leu        | Va I<br>840 | Ala        | Asn        | Leu        | Val        | Thr<br>845  | Ala        | Ser        | Lei        |
| Arg        | Arg<br>850 |            | Leu        | He         | Asp        | Arg<br>855 |             | Pro        | Glu        | Gly        | Ala<br>860 | Val         | Ala        | His        | Ala        |
| GIn<br>865 |            | Val        | lle        | Ser        | Asp<br>870 |            | Leu         | Cys        | Asn        | Arg<br>875 |            | Asp         | He         | Ser        | G11<br>880 |
| Leu        | Val        | lle        | Thr        | Lys<br>885 |            | Leu        | Thr         | Arg        | Ala<br>890 |            | Ser        | Asp         | Tyr        | Ala<br>895 | Gly        |
| Lys        | Gin        | Ala        | His<br>900 |            | Glu        | Leu        | Ala         | Glu<br>905 |            | ; Met      | Arg        | Lys         | Arg<br>910 | Asp        | Pro        |
| Gly        | ' Ser      | Ala<br>915 |            | Ser        | · Leu      | ı Gly      | 920         |            | ; Val      | Pro        | Tyr        | Va I<br>925 | He         | lie        | Se         |
| Ala        | 930        |            | Gly        | Val        | Ala        | 935        |             | Met        | : Lys      | Ser        | Glu<br>940 |             | Pro        | Leu        | Ph         |

<211> 6912

| Va I<br>945   | Leu         | Glu         | His         | Ser         | Leu<br>950  | Pro         | lle         | Asp         | Thr         | Gin<br>955  | Tyr         | Tyr                 | Leu         | Glu                  | GIn<br>960  |
|---------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|---------------------|-------------|----------------------|-------------|
| Gin           | Leu         | Ala         | Lys         | Pro<br>965  | Leu         | Leu         | Arg         | He          | Phe<br>970  | Glu         | Pro         | lle                 | Leu         | Gly<br>975           | Glu         |
| Gly           | Arg         | Ala         | Glu<br>980  | Ala         | Val         | Leu         | Leu         | Arg<br>985  | Gly         | Asp         | His         | Thr                 | Arg<br>990  | Cys                  | Lys         |
| Thr           | Val         | Leu<br>995  | Thr         | Gly         | Lys         |             | Gly<br>1000 | Gly         | Leu         | Leu         |             | Phe<br>1 <b>005</b> | Ala         | Lys                  | Arg         |
|               | Asn<br>1010 | Cys         | Cys         | lle         |             | Cys<br>1015 | Arg         | Thr         | Val         |             | Ser<br>1020 | His                 | Gin         | Gly                  | Ala         |
| Val<br>102    | -           | Glu         | Phe         | Cys         | Gin<br>1030 | Pro         | Arg         | Glu         |             | Glu<br>1035 | Leu         | Tyr                 | Gln         |                      | Glu<br>1040 |
| Val           | Ser         | His         |             | Asn<br>1045 | Ala         | Leu         | Glu         |             | Arg<br>1050 | Phe         | Ser         | Arg                 |             | Trp<br>1 <b>05</b> 5 | Thr         |
| Gin           | Cys         |             | Arg<br>1060 | Cys         | Gln         | Gly         |             | Leu<br>1065 | His         | Glu         | Asp         |                     | lle<br>1070 | Cys                  | Thr         |
| Ser           |             | Asp<br>1075 |             | Pro         | He          |             | Tyr<br>1080 | Met         | Arg         | Lys         |             | Va l<br>1085        | Arg         | Lys                  | Asp         |
|               | Glu<br>1090 |             | Gin         | Glu         |             | Leu<br>1095 |             | Arg         | Arg         |             | Gly<br>1100 | Pro                 | Pro         | Gly                  | Pro         |
| Glu<br>110    | Ala<br>5    | Trp         |             |             |             |             |             |             |             |             |             |                     |             |                      |             |
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#### <212> DNA

<213> Homo sapiens

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| gagccatcca gcgattcctg ctcgcctaca aggaggagcg ccgggggccc | acactcatcg | 4800 |
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<211> 2286

<212> PRT

<213> Homo sapiens

<400> 54

Met Ser Leu Arg Ser Gly Gly Arg Arg Arg Ala Asp Pro Gly Ala Asp 1 · 5 10 15

Gly Glu Ala Ser Arg Asp Asp Gly Ala Thr Ser Ser Val Ser Ala Leu 20 25 30

Lys Arg Leu Giu Arg Ser Gin Trp Thr Asp Lys Met Asp Leu Arg Phe 35 40 45

Gly Phe Glu Arg Leu Lys Glu Pro Gly Glu Lys Thr Gly Trp Leu IIe 50 55 60

Asn Met His Pro Thr Glu IIe Leu Asp Glu Asp Lys Arg Leu Gly Ser 65 70 75 80

Ala Val Asp Tyr Tyr Phe lle Gin Asp Asp Gly Ser Arg Phe Lys Val 85 90 95

Ala Leu Pro Tyr Lys Pro Tyr Phe Tyr lle Ala Thr Arg Lys Gly Cys
100 105 110

| Glu        | Arg        | Glu<br>115 | Val ·      | Ser        | Ser        | Phe<br>·    | Leu<br>120 | Ser        | Lys        | Lys        | Phe        | GIn<br>125 | Gly         | Lys        | lle         |
|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|-------------|
| Ala        | Lys<br>130 | Vai        | Glu        | Thr        | Val        | Pro<br>135  | Lys        | Glu        | Asp        | Leu        | Asp<br>140 | Leu        | Pro         | Asn        | His         |
| Leu<br>145 | Val        | Gly        | Leu        | Lys        | Arg<br>150 | <b>A</b> sn | Tyr        | He         | Arg        | Leu<br>155 | Ser        | Phe        | His         | Thr        | Val<br>160  |
| Glu        | Asp        | Leu        | Val        | Lys<br>165 | Val        | Arg         | Lys        | Glu        | lle<br>170 | Ser        | Pro        | Ala        | Val         | Lys<br>175 | Lys         |
| Asn        | Arg        | Glu        | GIn<br>180 | Asp        | His        | Ala         | Ser        | Asp<br>185 | Ala        | Tyr        | Thr        | Ala        | Leu<br>190  | Leu        | Ser         |
| Ser        | Val        | Leu<br>195 | Gln        | Arg        | Gly        | Gly         | Val<br>200 | He         | Thr        | Asp        | Glu        | Glu<br>205 | Glu         | Thr        | Ser         |
| Lys        | Lys<br>210 | lle        | Ala        | Asp        | Gln        | Leu<br>215  | Asp        | Asn        | lle        | Val        | Asp<br>220 | Met        | Arg         | Glu        | Tyr         |
| Asp<br>225 | Val        | Pro        | Tyr        | His        | lle<br>230 | Arg         | Leu        | Ser        | lle        | Asp<br>235 | Leu        | Lys        | ile         | His        | Va I<br>240 |
| Ala        | His        | Trp        | Tyr        | Asn<br>245 | Val        | Arg         | Tyr        | Arg        | Gly<br>250 | Asn        | Ala        | Phe        | Pro         | Val<br>255 | Glu         |
| lle        | Thr        | Arg        | Arg<br>260 | Asp        | Asp        | Leu         | Val        | Glu<br>265 | Arg        | Pro        | Asp        | Pro        | Va I<br>270 | Val        | Leu         |
| Ala        | Phe        | Asp<br>275 | He         | Glu        | Thr        | Thr         | Lys<br>280 | Leu        | Pro        | Leu        | Lys        | Phe<br>285 | Pro         | Asp        | Ala         |
| Glu        | Thr<br>290 | Asp        | Gin        | He         | Met        | Met<br>295  | lle        | Ser        | Tyr        | Met        | 11e<br>300 | Asp        | Gly         | Gln        | Gly         |

| Tyr<br>305 | Leu        | lle        | Thr        | Asn         | Arg<br>310 | Glu        | He         | Val        | Ser        | Glu<br>315 | Asp         | He         | Glu        | <b>A</b> sp | Phe<br>320 |
|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|-------------|------------|
| Glu        | Phe        | Thr        | Pro        | Lys<br>325  | Pro        | Glu        | Tyr        | Glu        | Gly<br>330 | Pro        | Phe         | Cys        | Val        | Phe<br>335  | Asn        |
| Glu        | Pro        | Asp        | Glu<br>340 | Ala         | His        | Leu        | lle        | GIn<br>345 | Arg        | Trp        | Phe         | Glu        | His<br>350 | Val         | GIn        |
| Glu        | Thr        | Lys<br>355 | Pro        | Thr         | He         | Met        | Val<br>360 | Thr        | Tyr        | Asn        | Gly         | Asp<br>365 | Phe        | Phe         | Asp        |
| Trp        | Pro<br>370 | Phe        | Val        | Glu         | Ala        | Arg<br>375 | Ala        | Ala        | Val        | His        | GIy<br>380  | Leu        | Ser        | Met         | Gin        |
| GIn<br>385 | Glu        | lle        | Gly        | Phe         | GIn<br>390 | Lys        | Asp        | Ser        | GIn        | Gly<br>395 | Glu         | Tyr        | Lys        | Ala         | Pro<br>400 |
| GIn        | Cys        | lle        | His        | Met<br>405  | Asp        | Cys        | Leu        | Arg        | Trp<br>410 | Val        | Lys         | Arg        | Asp        | Ser<br>415  | Tyr        |
| Leu        | Pro        | Val        | Gly<br>420 | Ser         | His        | Asn        | Leu        | Lys<br>425 | Ala        | Ala        | Ala         | Lys        | Ala<br>430 | Lys         | Leu        |
| Gly        |            | Asp<br>435 | Pro        | Val         | Glu        | Leu        | Asp<br>440 | Pro        | Glu        | Asp        | Met         | Cys<br>445 | Arg        | Met         | Ala        |
| Thr        | Glu<br>450 | Gln        | Pro        | ĠIn         | Thr        | Leu<br>455 | Ala        | Thr        | Tyr        | Ser        | Va I<br>460 | Ser        | Asp        | Ala         | Val        |
| Ala<br>465 | Thr        | Tyr        | Tyr        | Leu         | Tyr<br>470 | Met        | Lys        | Tyr        | Val        | His<br>475 | Pro         | Phe        | He         | Phe         | Ala<br>480 |
| Leu        | Cys        | Thr        | He         | 1 le<br>485 | Pro        | Met        | Glu        | Pro        | Asp<br>490 | Glu        | Val         | Leu        | Arg        | Lys<br>495  | Gly        |
| Ser        | Gly        | Thr        | Leu        | Cys         | Glu        | Ala        | Leu        | Leu        | Met        | Val        | Gin         | Ala        | Phe        | His         | Ala        |

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|             |            |             | 500         |            |            |            |                   | 505        |            |            |             |            | 510        |            |             |
|-------------|------------|-------------|-------------|------------|------------|------------|-------------------|------------|------------|------------|-------------|------------|------------|------------|-------------|
| Asn         | He         | l le<br>515 | Phe         | Pro        | Asn        | Lys        | Gln<br><b>520</b> | Glu        | Gln        | Glu        | Phe         | Asn<br>525 | Lys        | Leu        | Thr         |
| Asp         | Asp<br>530 | Gly         | His         | Val        | Leu        | Asp<br>535 | Ser               | Glu        | Thr        | Tyr        | Va I<br>540 | Gly        | Gly        | His        | Val         |
| Glu<br>545  | Ala        | Leu         | Glu         | Ser        | Gly<br>550 | Val        | Phe               | Arg        | Ser        | Asp<br>555 | lle         | Pro        | Cys        | Arg        | .Phe<br>560 |
| Arg         | Met        | Asn         | Pro         | Ala<br>565 | Ala        | Phe        | Asp               | Phe        | Leu<br>570 | Leu        | Gin         | Arg        | Val        | Glu<br>575 | Lys         |
| Thr         | Leu        | Arg         | His<br>580  | Ala        | Leu        | Glu        | Glu               | Glu<br>585 | Glu        | Lys        | Val         | Pro        | Val<br>590 | Glu        | Gln         |
| Val         | Thr        | Asn<br>595  | Phe         | Glu        | Glu        | Val        | Cys<br>600        | Asp        | Glu        | He         | Lys         | Ser<br>605 | Lys        | Leu        | Ala         |
| Ser         | Leu<br>610 | Lys         | <b>A</b> sp | Val        | Pro        | Ser<br>615 | Arg               | lle        | Glu        | Cys        | Pro<br>620  | Leu        | He         | Tyr        | His         |
| Leu<br>625  | Asp        | Val         | Gly         | Ala        | Met<br>630 | Tyr        | Pro               | Asn        | He         | lle<br>635 | Leu         | Thr        | Asn        | Arg        | Leu<br>640  |
| Gln         | Pro        | Ser         | Ala         | Met<br>645 | Val        | Asp        | Glu               | Ala        | Thr<br>650 | Cys        | Ala         | Ala        | Cys        | Asp<br>655 | Phe         |
| <b>A</b> sn | Lys        | Pro         | Gly<br>660  | Ala        | Asn        | Cys        | Gln               | Arg<br>665 | Lys        | Met        | Ala         | Trp        | GIn<br>670 | Trp        | Arg         |
| Gly         | Glu        | Phe<br>675  | Met         | Pro        | Ala        | Ser        | Arg<br>680        | Ser        | Glu        | Tyr        | His         | Arg<br>685 | He         | GIn        | His         |
| Gln         | Leu<br>690 | Glu         | Ser         | Glu        | Lys        | Phe<br>695 | Pro               | Pro        | Leu        | Phe        | Pro<br>700  | Glu        | Gly        | Pro        | Ala         |

| Arg<br>705 | Ala        | Phe         | His        | Glu        | 110        | Ser        | Arg        | Glu        | GIU         | 715        | Ala        | Lys        | lyr        | Giu        | 120        |
|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|
| Arg        | Arg        | Leu         | Ala        | Asp<br>725 | Tyr        | Cys        | Arg        | Lys        | Ala<br>730  | Tyr        | Lys        | Lys        | lle        | His<br>735 | He         |
| Thr        | Lys        | Val         | Glu<br>740 | Glu        | Arg        | Leu        | Thr        | Thr<br>745 | He          | Cys        | GIn        | Arg        | Glu<br>750 | Asn        | Ser        |
| Phe        | Tyr        | Va I<br>755 | Asp        | Thr        | Val        | Arg        | Ala<br>760 | Phe        | Arg         | Asp        | Arg        | Arg<br>765 | Tyr        | Glu        | Phe        |
| Lys        | Gly<br>770 | Leu         | His        | Lys        | Val        | Trp<br>775 | Lys        | Lys        | Lys         | Leu        | Ser<br>780 | Ala        | Ala        | Val        | Glu        |
| Va!<br>785 | Gly        | Asp         | Ala        | Ala        | Glu<br>790 | Val        | Lys        | Arg        | Cys         | Lys<br>795 | Asn        | Met        | Glu        | Val        | Leu<br>800 |
| Tyr        | Asp        | Ser         | Leu        | G1n<br>805 | Leu        | Ala        | His        | Lys        | Cys<br>810  | lle        | Leu        | Asn        | Ser        | Phe<br>815 | Tyr        |
| Gly        | Tyr        | Val         | Met<br>820 | Arg        | Lys        | Gly        | Ala        | Arg<br>825 | Trp         | Tyr        | Ser        | Met        | Glu<br>830 | Met        | Ala        |
| Gly        | He         | Va I<br>835 | Cys        | Phe        | Thr        | Gly        | Ala<br>840 | Asn        | He          | lle        | Thr        | Gln<br>845 | Ala        | Arg        | Glu        |
| Leu        | lle<br>850 | Glu         | Gln        | lle        | Gly        | Arg<br>855 | Pro        | Leu        | Glu         | Leu        | Asp<br>860 | Thr        | Asp        | Gly        | lle        |
| Trp<br>865 | Cys        | Val         | Leu        | Pro        | Asn<br>870 | Ser        | Phe        | Pro        | Glu         | Asn<br>875 | Phe        | Val        | Phe        | Lys        | Thr<br>880 |
| Thr        | Asn        | Val         | Lys        | Lys<br>885 | Pro        | Lys        | Val        | Thr        | l le<br>890 | Ser        | Tyr        | Pro        | Gly        | Ala<br>895 | Met        |

| Leu         | Asn         | lle         | Met<br>900  | Val         | Lys         | Glu         | Gly                 | Phe<br>905  | Thr         | Asn                | Asp         | Gln         | Tyr<br>910  | Gin         | Glu         |
|-------------|-------------|-------------|-------------|-------------|-------------|-------------|---------------------|-------------|-------------|--------------------|-------------|-------------|-------------|-------------|-------------|
| Leu         | Ala         | Glu<br>915  | Pro         | Ser         | Ser         | Leu         | Thr<br>920          |             | Vai         | Thr                | Arg         | Ser<br>925  | Glu         | Asn         | Ser         |
| lle         | Phe<br>930  | Phe         | Glu         | Val         | Asp         | Gly<br>935  | Pro                 | Tyr         | Leu         | Ala                | Met<br>940  | He          | Leu         | Pro         | Ala         |
| Ser<br>945  | Lys         | Glu         | Glu         | Gly         | Lys<br>950  | Lys         | Leu                 | Lys         | Lys         | <b>A</b> rg<br>955 | Tyr         | Ala         | Val         | Phe         | Asn<br>960  |
| Glu         | Asp         | Gly         | Ser         | Leu<br>965  | Ala         | Glu         | Leu                 | Lys         | Gly<br>970  | Phe                | Glu         | Val         | Lys         | Arg<br>975  | Arg         |
| Gly         | Glu         | Leu         | GIn<br>980  | Leu         | He          | Lys         | He                  | Phe<br>985  | Gin         | Ser                | Ser         | Val         | Phe<br>990  | Glu         | Ala         |
| Phe         | Leu         | Lys<br>995  | Gly         | Ser         | Thr         |             | Glu<br>1 <b>000</b> | Glu         | Val         | Tyr                |             | Ser<br>1005 | Val         | Ala         | Lys         |
|             | Ala<br>1010 | Asp         | Tyr         | Trp         |             | Asp<br>1015 | Val                 | Leu         | Tyr         |                    | Lys<br>1020 | Ala         | Ala         | Asn         | Met         |
| Pro<br>1025 |             | Ser         | Glu         |             | Phe<br>1030 | Glu         | Leu                 | He          |             | Glu<br>1035        | Asn         | Arg         | Ser         |             | Ser<br>1040 |
| Arg         | Lys         | Leu         |             | Asp<br>1045 | Tyr         | Gly         | Glu                 |             | Lys<br>1050 | Ser                | Thr         | Ser         |             | Ser<br>1055 | Thr         |
| Ala         | Lys         |             | Leu<br>1060 | Ala         | Glu         | Phe         |                     | Gly<br>1065 | Asp         | Gin                | Met         | Val         | Lys<br>1070 | Asp         | Ala         |
| Gly         |             | Ser<br>1075 | Cys         | Arg         | Tyr         |             | l le<br>1080        | Ser         | Arg         | Lys                |             | Glu<br>1085 | Gly         | Ser         | Pro         |
| Val         | Thr         | Glu         | Arg         | Ala         | He          | Pro         | Leu                 | Ala         | He          | Phe                | Gln         | Ala         | Glu         | Pro         | Thr         |

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| 1090              |                      | 1095                            | 11                    | 00                          |
|-------------------|----------------------|---------------------------------|-----------------------|-----------------------------|
| Val Arg I<br>1105 |                      | _eu Arg Lys<br>110              | Trp Leu Lys S<br>1115 | Ser Ser Ser Leu Gln<br>1120 |
| Asp Phe /         | Asp lie Arg /        |                                 | Asp Trp Asp 1         | Tyr lle Glu Arg<br>1135     |
| Leu Gly S         | Ser Ala Ile (        |                                 | lle Thr lle F<br>145  | Pro Ala Ala Leu Gin<br>1150 |
|                   | Lys Asn Pro 1<br>155 | Val Pro Arg<br>1160             | Val Lys His F         | Pro Asp Trp Leu His<br>1165 |
| Lys Lys I<br>1170 | Leu Leu Glu          | Lys <b>A</b> sn Asp<br>1175     | •                     | âin Lys Lys lle Ser<br>180  |
| Glu Leu I<br>1185 |                      | Glu Gly Arg<br>190 <sub>.</sub> | Arg Gln Val 1         | Thr Met Ala Glu Ala<br>1200 |
| Ser Glu           | Asp Ser Pro<br>1205  | Arg Pro Ser                     | Ala Pro Asp I<br>1210 | Met Glu Asp Phe Gly<br>1215 |
| Leu Val           | Lys Leu Pro<br>1220  |                                 | Ala Pro Val           | Thr Vai Lys Arg Lys<br>1230 |
|                   | Leu Trp Glu<br>235   | Ser Gin Giu<br>1240             | Glu Ser Gln /         | Asp Leu Thr Pro Thr<br>1245 |
| Val Pro<br>1250   | Trp Gln Glu          | lle Leu Gly<br>1255             |                       | Ala Leu Gly Thr Ser<br>260  |
| Gin Glu<br>1265   | ·                    | Val Trp Leu<br>270              | Arg Phe His 1<br>1275 | Lys Lys Trp Gin<br>1280     |
| Leu Gin           | Ala Arg Gin<br>1285  | Arg Leu Ala                     | Arg Arg Lys 1         | Arg Gln Arg Leu Glu<br>1295 |

| Ser Ala   | Glu  | Gly  | Val  | Leu  | Arg   | Pro         | Gly      | Ala  | He   | Arg      | Asp        | Gly   | Pro  | Ala  |
|-----------|------|------|------|------|-------|-------------|----------|------|------|----------|------------|-------|------|------|
|           | 1    | 1300 |      |      |       | 1           | 305      |      |      |          | 1          | 310   |      |      |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Thr Gly   | Leu  | Gly  | Ser  | Phe  | Leu   | Arg         | Arg      | Thr  | Ala  | Arg      | Ser        | He    | Leu  | Asp  |
| •         | 1315 |      |      |      | 1     | 1320        |          |      |      | •        | 1325       |       |      |      |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Leu Pro   | Trp  | Gin  | He   | Val  | Gln   | He          | Ser      | Glu  | Thr  | Ser      | Gln        | Ala   | Gly  | Leu  |
| 1330      | •    |      |      |      | 1335  |             |          |      |      | 1340     |            |       |      |      |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Phe Arg   | Leu  | Tro  | Ala  | Leu  | Val   | Glv         | Ser      | Asp  | Leu  | His      | Cvs        | He    | Arg  | Leu  |
| 1345      |      | 6    |      | 350  |       | ,           | <b>.</b> |      | 1355 | •••      | -,-        | • • • |      | 1360 |
| 10-10     |      |      | ,    | 000  |       |             |          |      | 1000 |          |            |       |      | .000 |
| Ser lle   | Pro  | Ara  | Val  | Dha  | Tvr   | Val         | Aen      | Gln  | Ara  | Val      | Δla        | l ve  | Δla  | Glu  |
| 361 116   | 110  |      | 1365 | 1110 | 1 91  | <b>V</b> 41 |          | 1370 | ЛΙБ  | <b>V</b> | πια        |       | 1375 | uiu  |
|           |      |      | 1300 |      |       |             |          | 1370 |      |          |            |       | 1070 |      |
| Clar Clar | Ala  | Cam  | T    | A    | 1     | Va I        | A        | Awa  | Va I | Lau      | Dro        | A-~   | Sor. | Aon  |
| Glu Gly   |      |      | ıyr  | Arg  | Lys   |             |          | Arg  | Vai  | Leu      |            |       | Ser  | ASH  |
|           |      | 1380 |      |      |       |             | 1385     |      |      |          |            | 1390  |      |      |
|           | _    |      |      | _    | •     | _           | _        |      |      | ٥,       |            |       | _    | ٥.   |
| Met Val   |      |      | Leu  | lyr  |       |             | Ser      | Vai  | Pro  |          |            | Met   | ıyr  | Gin  |
| •         | 1395 |      |      |      | •     | 1400        |          |      |      | •        | 1405       |       |      |      |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Glu His   | He   | Asn  | Glu  | lle  | Asn   | Ala         | Glu      | Leu  | Ser  | Ala      | Pro        | Asp   | He   | Glu  |
| 1410      |      |      |      | •    | 1415  | •           |          |      | •    | 1420     |            |       |      |      |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Gly Vai   | Tyr  | Glu  | Thr  | Gin  | Vai   | Pro         | Leu      | Leu  | Phe  | Arg      | Ala        | Leu   | Val  | His  |
| 1425      |      |      |      | 1430 |       |             |          | •    | 1435 |          |            |       | 1    | 1440 |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Leu Gly   | Cys  | Val  | Cys  | Val  | Val   | Asn         | Lys      | Gln  | Leu  | Val      | Arg        | His   | Leu  | Ser  |
|           |      | •    | 1445 |      |       |             |          | 1450 |      |          |            |       | 1455 |      |
|           |      |      |      |      |       |             |          |      |      | •        |            |       |      |      |
| Gly Trp   | Glu  | Ala  | Glu  | Thr  | Phe   | Ala         | Leu      | Glu  | His  | Leu      | Glu        | Met   | Arg  | Ser  |
|           |      | 1460 |      |      |       |             | 1465     |      |      |          |            | 1470  | _    |      |
|           |      |      |      |      |       |             |          |      |      |          |            |       |      |      |
| Leu Ala   | Gln  | Pho  | Ser  | Tvr  | יום ו | Glii        | Pro      | Glv  | Ser  | ماا      | Ara        | Hie   | ما   | Tvr  |
|           |      |      | OGI  | ı yı |       | 1480        | . 10     | uly  | 061  |          | ль<br>1485 | 1110  | , 10 |      |
|           | 1475 |      |      |      |       | 146U        |          |      |      |          | 1400       |       |      |      |

| Leu Tyr His<br>1490 |                     | Ala His Lys<br>1495 | S Ala Leu Phe<br>1500 | Gly lle Phe lle           |
|---------------------|---------------------|---------------------|-----------------------|---------------------------|
| Pro Ser Glh<br>1505 | Arg Arg Ala<br>1510 |                     | e Val Leu Asp<br>1515 | Thr Val Arg Ser<br>1520   |
| Asn Gin Met         | Pro Ser Leu<br>1525 | Gly Ala Le          | ı Tyr Ser Ala<br>1530 | Glu His Gly Leu<br>1535   |
|                     | Lys Val Gly<br>1540 | Pro Glu Let         |                       | Pro Lys His Thr<br>. 1550 |
| Phe Glu Val<br>1555 | Arg Ala Glu         | Thr Asp Let         |                       | Cys Arg Ala Ile<br>1565   |
| Gln Arg Phe<br>1570 |                     | Tyr Lys Gli<br>1575 | u Glu Arg Arg<br>1580 | Gly Pro Thr Leu           |
| lle Ala Val<br>1585 | Gin Ser Ser<br>1590 | Trp Glu Le          | ı Lys Arg Leu<br>1595 | Ala Ser Glu Ile<br>1600   |
| Pro Val Leu         | Glu Glu Phe<br>1605 | Pro Leu Va          | l Pro IIe Cys<br>1610 | Val Ala Asp. Lys<br>1615  |
|                     | Gly Val Leu<br>1620 | Asp Trp Gli<br>162  |                       | Ala Arg Arg Met<br>1630   |
| lle Arg His<br>1635 | Tyr Leu Asn         | Leu Asp Thi<br>1640 |                       | Gin Ala Phe Giu<br>1645   |
| Met Ser Arg         |                     | lle Pro lle<br>1655 | e Gly Asn Leu<br>1660 | Pro Glu Asp Ile           |
| Ser Thr Phe<br>1665 | Gly Ser Asp<br>1670 | Leu Phe Pho         | e Ala Arg His<br>1675 | Leu Gin Arg His<br>1680   |
| Asn His Leu         | Leu Trp Leu         | Ser Pro Thi         | - Ala Arg Pro         | Asp Leu Gly Gly           |

|                         | 1685              | 169                    | 90                     | 1695                |
|-------------------------|-------------------|------------------------|------------------------|---------------------|
| Lys Glu Ala Asp<br>1700 | Asp Asn C         | Cys Leu Val Mi<br>1705 | •                      | Asp Gln Ala<br>1710 |
| Thr Val Glu lle<br>1715 | Asn Ser S         | Ser Gly Cys Ty<br>1720 | yr Ser Thr Val<br>1725 |                     |
| Leu Asp Leu Gin<br>1730 |                   | Ala Val Asn Ti<br>735  | hr lle Leu Gin<br>1740 | Ser His His         |
| Val Asn Asp Met<br>1745 | Glu Gly A<br>1750 | Ala Asp Ser M          | et Gly lle Ser<br>1755 | Phe Asp Val         |
| lle Gin Gin Ala         | Ser Leu 6<br>1765 | Glu Asp Met I<br>17    |                        | Gin Ala Ala<br>1775 |
| Ser Ala Pro Ala<br>1780 | Ser Tyr A         | Asp Glu Thr A<br>1785  |                        | Asn Thr Phe<br>1790 |
| Arg lie Leu Lys<br>1795 | Ser Met V         | /al Val Gly Ti<br>1800 | rp Val Lys Glu<br>1805 | lle Thr Gin         |
| Tyr His Asn lie<br>1810 |                   | Asp Asn Gin Va<br>315  | al Met His Phe<br>1820 | Tyr Arg Trp         |
| Leu Arg Ser Pro<br>1825 | Ser Ser L<br>1830 | eu Leu His A           | sp Pro Ala Leu<br>1835 | His Arg Thr<br>1840 |
| Leu His Asn Met         | Met Lys L<br>1845 | ys Leu Phe Lo          | •                      | Ala Glu Phe<br>1855 |
| Lys Arg Leu Gly<br>1860 | Ser Ser V         | /al lle Tyr A<br>1865  |                        | Arg lie lie<br>1870 |
| Leu Cys Thr Lys<br>1875 | Lys Arg A         | Arg Val Glu As<br>1880 | sp Ala Ile Ala<br>1885 | Tyr Val Glu         |

|             | l le<br>1890 | Thr         | Ser         | Ser          |             | His<br>1895 | Ser         | Lys          | Glu         |             | Phe<br>1900 | His         | Ser         | Leu         | Inr         |
|-------------|--------------|-------------|-------------|--------------|-------------|-------------|-------------|--------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|
| l le<br>190 | Ser<br>5     | Phe         | Ser         |              | Cys<br>1910 | Trp         | Glu         | Phe          |             | Leu<br>1915 | Trp         | Met         | Asp         |             | Ser<br>1920 |
| Asn         | Tyr          | Gly         |             | l le<br>1925 | Lys         | Gly         | Lys         |              | Ser<br>1930 | Ser         | Arg         | He          |             | Cys<br>1935 | Gly         |
| Leu         | GIn          | -           | Ser<br>1940 | Gln          | Lys         | Ala         |             | Gly<br>1945  | Ala         | Glu         | Asp         |             | GIn<br>1950 | Glu         | Asn         |
| Glu         | Asp          | Asp<br>1955 | Glu         | Glu          | Glu         |             | Asp<br>1960 | Gly          | Glu         | Glu         |             | Glu<br>1965 | Glu         | Ala         | Glu         |
|             | Ser<br>1970  |             | Val         | Glu          |             | Leu<br>1975 | Leu         | Glu          | Asn         |             | Trp<br>1980 | Asn         | He          | Leu         | Gln         |
| Phe<br>198  | Leu<br>5     | Pro         | Gln         |              | Ala<br>1990 | Ser         | Cys         | GIn          |             | Tyr<br>1995 | Phe         | Leu         | Met         |             | Val<br>2000 |
| Ser         | Ala          | Tyr         |             | Val<br>2005  | Ala         | Val         | Tyr         |              | Cys<br>2010 | Met         | Lys         | Asp         |             | Leu<br>2015 | Arg         |
| Arg         | Ser          |             | Pro<br>2020 | Gly          | Ser         | Thr         |             | Va I<br>2025 | Arg         | Arg         | Arg         |             | Ala<br>2030 | Ser         | Gin         |
| Leu         | Ser          | GIn<br>2035 | Glu         | Ala          | Glu         |             | Ala<br>2040 | Val          | Gly         | Ala         |             | Pro<br>2045 | Gly         | Met         | lle         |
|             | Phe<br>2050  |             | Gln         | Asp          |             | Val<br>2055 | Ala         | <b>As</b> n  |             |             | Thr<br>2060 | Gin         | Ser         | Phe         | Phe         |
| Thr<br>206  | ile<br>5     | Thr         | Gln         | -            | lle<br>2070 | Gln         | Lys         | Lys          |             | Thr<br>2075 | Gly         | Ser         | Arg         |             | Ser<br>2080 |

| Thr        | Glu         | Leu         |              | Glu<br>2085   | Met         | Phe         | Pro         |               | Leu<br>2090 | Pro         | Gly         | Ser         |               | Leu<br>2095    | Leu         |
|------------|-------------|-------------|--------------|---------------|-------------|-------------|-------------|---------------|-------------|-------------|-------------|-------------|---------------|----------------|-------------|
| Leu        | Asn         |             | Pro<br>2100  | Ala           | Leu         | Glu         |             | lle<br>2105   | Lys         | Tyr         | Val         |             | Lys<br>2110   | Val            | Leu         |
| Ser        |             | Asp<br>2115 |              | Asn           | lle         |             | Asn<br>2120 | Gln           | Vai         | Asn         |             | Leu<br>2125 |               | Arg            | Asp         |
|            | Leu<br>2130 |             | Leu          | Val           |             | Val<br>2135 | Gly         | Glu           | Phe         |             | Glu<br>2140 |             | Ala           | Gln            | Phe         |
| Arg<br>214 |             | Pro         | Cys          | Arg           | Ser<br>2150 | Tyr         | Val         | Leu           |             | Glu<br>2155 | Val         | He          | Cys           |                | Ser<br>2160 |
| Cys        | Asn         | Phe         |              | Arg<br>2165   |             | Leu         | Asp         |               | Cys<br>2170 |             | Asp         | Ser         |               | Phe<br>2175    | Ser         |
| Glu        | Asp         | _           | Ala<br>2180  | Val           | Leu         | Pro         |             | Trp<br>2185   |             | Cys         | Ser         |             | Cys<br>2190   |                | Ala         |
| Pro        | -           | Asp<br>2195 |              | · Ser         | Ala         |             | Glu<br>2200 |               | Thr         | Leu         | Val         | Glu<br>2205 |               | Leu            | GIn         |
|            | Lys<br>2210 |             | ı Met        | : Ala         |             | Thr<br>2215 |             | Gln           | <b>A</b> sp |             | Val<br>2220 |             | Leu           | Lys            | Cys         |
| Arg<br>222 | -           | <b>Va</b> l | Lys          | s-Glu         | Thr<br>2230 |             | Met         | : Pro         |             | Tyr<br>2235 |             | Thr         | · Cys         |                | Gly<br>2240 |
| Asp        | Phe         | e Ala       | a Lei        | ı Thr<br>2245 | _           | His         | Thr         | Glr           | Val<br>2250 |             | Met         | : Glu       | ı Gin         | 1   le<br>2255 |             |
| l le       | : Phe       | e Are       | 2260<br>2260 | 1     E       | Ala         | Gln         |             | : Tyr<br>2265 |             | Met         | : Ser       | - Tyr       | · Leu<br>2270 |                | Glu         |

Thr Leu Glu Trp Leu Leu Gln Lys Asn Pro Gln Leu Gly His

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#### 102/236

2275 2280 2285

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| gtcaaggtcg | tgtctcagct | gctgcgccag | gccatgcgcc | aggggctgct | gatgcctgtg | 1740 |
|------------|------------|------------|------------|------------|------------|------|
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| tactatgacg | tccccattgc | caccctggac | ttctcctcct | tgtacccatc | catcatgatg | 1860 |
| gcccataatc | tgtgctacac | cacgctgctc | cgacctgggg | ctgcccagaa | gctgggcctt | 1920 |
| aaaccagatg | agttcatcaa | gacacccact | ggggatgagt | ttgtgaagtc | atctgtacgg | 1980 |
| aagggcctcc | tgccccagat | cctggagaat | ctgctgagtg | cccgcaagag | ggccaaggct | 2040 |
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| tacggtgaca | cggactctgt | gatgtgccgg | tttggcgtct | cctctgtggc | tgaagcaatg | 2340 |
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| gagttcgaga | aggtttactt | cccatacctg | ctcatcagca | agaagcgcta | tgctggcctg | 2460 |
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| tgcaaccgca | tagacatctc | ccagctggtc | atcaccaaag | agttgacccg | cgcagcagca | 2700 |
| gactatgctg | gcaagcaggc | tcacgtggag | ctggctgaga | ggatgaggaa | gcgcgacccc | 2760 |
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|            |            |            |            | •          |            |      |
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| gaacggctgc | tgcagcgctt | tggaccgccc | ggccctgagg | cctggtgacc | tgacacggga | 3360 |
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<212> PRT

<213> Mus musculus

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|------------|------------|------------|------------|------------|------------|-------------|-------------|-------------|-------------|------------|------------|------------|------------|------------|------------|
| Met<br>1   | Asp        | Cys        | Lys        | Arg<br>5   | Arg        | Gln         | Gly         | Pro         | Gly<br>10   | Pro        | Gly        | Val        | Pro        | Pro<br>15  | Lys        |
| Arg        | Ala        | Arg        | Gly<br>20  | His        | Leu        | Trp         | Asp         | Glu<br>25   | Asp         | Glu        | Pro        | Ser        | Pro<br>30  | Ser        | Gln        |
| Phe        | Glu        | Ala<br>35  | Asn        | Leu        | Ala        | Leu         | Leu<br>40   | Glu         | Glu         | He         | Glu        | Ala<br>45  | Glu        | Asn        | Arg        |
| Leu        | GIn<br>50  | Glu        | Ala        | Glu        | Glu        | Glu<br>55   | Leu         | Gin         | Leu         | Pro        | Pro<br>60  | Glu        | Gly        | Thr        | Val        |
| Gly<br>65  | Gly        | GIn        | Phe        | Ser        | Thr<br>70  | Ala         | <b>A</b> sp | He          | Asp         | Pro<br>75  | Arg        | Trp        | Arg        | Arg        | Pro<br>80  |
| Thr        | Leu        | Arg        | Ala        | Leu<br>85  | Asp        | Pro         | Ser         | Thr         | Glu<br>90   | Pro        | Leu        | lle        | Phe        | GIn<br>95  | Gln        |
| Leu        | Glu        | lle        | Asp<br>100 | His        | Tyr        | Val         | Gly         | Ser<br>105  | Ala         | Pro        | Pro        | Leu        | Pro<br>110 | Glu        | Arg        |
| Pro        | Leu        | Pro<br>115 | Ser        | Arg        | Asn        | Ser         | Va l<br>120 | Pro         | lle         | Leu        | Arg        | Ala<br>125 | Phe        | Gly        | Val        |
| Thr        | Asp<br>130 |            | Gly        | Phe        | Ser        | Va I<br>135 | Cys         | Cys         | His         | lle        | Gln<br>140 | Gly        | Phe        | Ala        | Pro        |
| Tyr<br>145 | Phe        | Tyr        | Thr        | Pro        | Ala<br>150 | Pro         | Pro         | Gly         | Phe         | Gly<br>155 | Ala        | Glu        | His        | Leu        | Ser<br>160 |
| Glu        | Leu        | Gln        | Gln        | Glu<br>165 |            | Asn         | Ala         | Ala         | l le<br>170 | Ser        | Arg        | Asp        | Gln        | Arg<br>175 | Gly        |
| Gly        | Lys        | Glu        | Leu<br>180 | Ser        | Gly        | Pro         | Ala         | Va I<br>185 | Leu         | Ala        | He         | Glu        | Leu<br>190 | Cys        | Ser        |

| Arg         | Glu         | Ser<br>195 | Met        | Phe        | Gly         | Tyr        | His<br>200 | Gly        | His        | Gly         | Pro        | Ser<br>205 | Pro        | Phe         | Leu        |
|-------------|-------------|------------|------------|------------|-------------|------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|
| Arg         | 1 le<br>210 | Thr        | Leu        | Ala        | Leu         | Pro<br>215 | Arg        | Leu        | Met        | Ala         | Pro<br>220 | Ala        | Arg        | Arg         | Leu        |
| Leu<br>225  | Glu         | Gln        | Gly        | Val        | Arg<br>230  | Val        | Pro        | Gly        | Leu        | Gly<br>235  | Thr        | Pro        | Ser        | Phe         | Ala<br>240 |
| Pro         | Tyr         | Glu        | Ala        | Asn<br>245 | Val         | Asp        | Phe        | Glu        | lle<br>250 | Arg         | Phe        | Met        | Val        | Asp<br>255  | Ala        |
| Asp         | He          | Val        | Gly<br>260 | Cys        | Asn         | Trp        | Leu        | Glu<br>265 | Leu        | Pro         | Ala        | Gly        | Lys<br>270 | Tyr         | Val        |
| Arg         | Arg         | Ala<br>275 | Glu        | Lys        | Lys         | Ala        | Thr<br>280 | Leu        | Cys        | Gin         | Leu        | Glu<br>285 | Val        | Asp         | Val        |
| Leu         | Trp<br>290  | Ser        | Asp        | Val        | He          | Ser<br>295 | His        | Pro        | Pro        | Glu         | Gly<br>300 | Gln        | Trp        | Gln         | Arg        |
| l le<br>305 | Ala         | Pro        | Leu        | Arg        | Va I<br>310 | Leu        | Ser        | Phe        | Asp        | 1 le<br>315 | Glu        | Ċys        | Ala        | Gly         | Arg<br>320 |
| Lys         | Gly         | He         | Phe        | Pro<br>325 | Glu         | Pro        | Glu        | Arg        | Asp<br>330 | Pro         | Val        | He         | Gln        | l le<br>335 |            |
| Ser         | Leu         | Gly        | Leu<br>340 |            | Trp         | Gly        | Glu        | Pro<br>345 | Glu        | Pro         | Phe        | Leu        | Arg<br>350 | Leu         | Ala        |
| Leu         | Thr         | Leu<br>355 | Arg        | Pro        | Cys         | Ala        | Pro<br>360 | lle        | Leu        | Gly         | Ala        | Lys<br>365 | Val        | Gln         | Ser        |
| Tyr         | G1u<br>370  | Arg        | Glu        | Glu        | Asp         | Leu<br>375 |            | GIn        | Ala        | Trp         | Ala<br>380 | Asp        | Phe        | He          | Leu        |
| Ala         | Met         | Asp        | Pro        | Asp        | Val         | lle        | Thr        | Gly        | Tyr        | Asn         | He         | Gln        | Asn        | Phe         | Asp        |

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| 385        |             |            |            |            | 390        |            |            |            |            | 395        |                  |            |            |            | 400        |
|------------|-------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|
| Leu        | Pro         | Tyr        |            | lle<br>405 | Ser        | Arg        | Ala        | Gin        | Ala<br>410 | Leu        | Lys <sub>.</sub> | Val        |            | Arg<br>415 | Phe        |
| Pro        | Phe         | Leu        | Gly<br>420 | Arg        | Val        | Thr        | Gly        | Leu<br>425 | Arg        | Ser        | Asn              | lle        | Arg<br>430 | Asp        | Ser        |
| Ser        | Phe         | GIn<br>435 | Ser        | Arg        | Gln        | Val        | Gly<br>440 | Arg        | Arg        | Asp        | Ser              | Lys<br>445 | Val        | He         | Ser        |
| Met        | Va I<br>450 | Gly        | Arg        | Val        | Gln        | Met<br>455 | Asp        | Met        | Leu        | GIn        | Val<br>460       | Leu        | Leu        | Arg        | Glu        |
| His<br>465 | Lys         | Leu        | Arg        | Ser        | Tyr<br>470 | Thr        | Leu        | Asn        | Ala        | Val<br>475 | Ser              | Phe        | His        | Phe        | Leu<br>480 |
| Gly        | Glu         | GIn        | Lys        | Glu<br>485 |            | Val        | Gln        | His        | Ser<br>490 | l le       | lle              | Thr        | Asp        | Leu<br>495 | Gin        |
| Asn        | Gly         | Asn        | Glu<br>500 | Gln        | Thr        | Arg        | Arg        | Arg<br>505 | Leu        | Ala        | Val              | Tyr        | Cys<br>510 | Leu        | Lys        |
| Asp        | Ala         | Phe<br>515 |            | Pro        | Leu        | Arg        | Leu<br>520 |            | Glu        | Arg        | Leu              | Met<br>525 | Val        | Leu        | Val        |
| Asn        | Asn<br>530  |            | Glu        | Met        | Ala        | Arg<br>535 |            | Thr        | Gly        | Val        | Pro<br>540       | Leu        | Gly        | Tyr        | Leu        |
| Leu<br>545 |             | Arg        | Gly        | Gln        | GIn<br>550 |            | Lys        | Val        | Val        | Ser<br>555 |                  | Leu        | Leu        | Arg        | GIn<br>560 |
| Ala        | Met         | Arg        | Gin        | Gly<br>565 |            | Leu        | Met        | Pro        | Val<br>570 |            | Lys              | Thr        | Glu        | Gly<br>575 | Gly        |
| Glu        | Asp         | Tyr        | Thr<br>580 |            | Ala        | Thr        | Vai        | 11e<br>585 |            | Pro        | Leu              | Lys        | Gly<br>590 |            | Tyr        |

| ASP        | Val        | 595        | 116        | AIA         | ınr        | Leu        | 600        | rne        | ser        | ser        |             | 605        | Pro        | Ser        | 116        |
|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|
| Met        | Met<br>610 | Ala        | His        | Asn         | Leu        | Cys<br>615 | Tyr        | Thr        | Thr        | Leu        | Leu<br>620  | Arg        | Pro        | Gly        | Ala        |
| Ala<br>625 | Gin        | Lys        | Leu        | Gly         | Leu<br>630 | Lys        | Pro        | Asp        | Glu        | Phe<br>635 | lle         | Lys        | Thr        | Pro<br>·   | Thr<br>640 |
| Gly        | Asp        | Glu        | Phe        | Va I<br>645 | Lys        | Ser        | Ser        | Val        | Arg<br>650 | Lys        | Gly         | Leu        | Leu        | Pro<br>655 | Gin        |
| He         | Leu        | Glu        | Asn<br>660 | Leu         | Leu        | Ser        | Ala        | Arg<br>665 | Lys        | Arg        | Ala         | Lys        | Ala<br>670 | Glu        | Leu        |
| Ala        | Gln        | Glu<br>675 |            | Asp         | Pro        | Leu        | Arg<br>680 | Arg        | Gln        | Val        | Leu         | Asp<br>685 | Gly        | Arg        | Gin        |
| Leu        | Ala<br>690 | Leu        | Lys        | Val         | Ser        | Ala<br>695 |            | Ser        | Val        | Tyr        | Gly<br>700  | Phe        | Thr        | Gly        | Ala        |
| GIn<br>705 |            | Gly        | Lys        | Leu         | Pro<br>710 | Cys        | Leu        | Glu        | lle        | Ser<br>715 |             | Ser        | Val        | Thr        | Gly<br>720 |
| Phe        | Gly        | Arg        | Gln        | Met<br>725  |            | Glu        | Lys        | Thr        | Lys<br>730 |            | Leu         | Val        | Glu        | Ser<br>735 | Lys        |
| Tyr        | Thr        | Val        | Glu<br>740 |             | Gly        | Tyr        | Asp        | Ala<br>745 |            | Ala        | Lys         | Val        | Val<br>750 |            | Gly        |
| Asp        | Thr        | Asp<br>755 |            | Val         | Met        | Cys        | Arg<br>760 |            | Gly        | Val        | Ser         | Ser<br>765 |            | Ala        | Glu        |
| Ala        | Met<br>770 | Ser        | Leu        | Gly         | Arg        | Glu<br>775 |            | Ala        | Asn        | Тгр        | Va I<br>780 |            | Ser        | His        | Phe        |

| Pro<br>785  | Ser        | Pro        | lle        | Arg        | Leu<br>790 | Glu        | Phe        | Glu        | Lys        | Va I<br>795 | Tyr        | Phe         | Pro         | Tyr        | Leu<br>800 |
|-------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|-------------|-------------|------------|------------|
| Leu         | ile        | Ser        | Lys        | Lys<br>805 | Arg        | Tyr        | Ala        | Gly        | Leu<br>810 | Leu         | Phe        | Ser         | Ser         | Arg<br>815 | Ser        |
| Asp         | Ala        | His        | Asp<br>820 | Lys        | Met        | Asp        | Cys        | Lys<br>825 | Gly        | Leu         | Glu        | Ala         | Va I<br>830 | Arg        | Arg        |
| <b>A</b> sp | Asn        | Cys<br>835 | Pro        | Leu        | Val        | Ala        | Asn<br>840 | Leu        | Val        | Thr         | Ser        | Ser<br>845  | Leu         | Arg        | Arg        |
| lle         | Leu<br>850 | Val        | Asp        | Arg        | Asp        | Pro<br>855 | Asp        | Gly        | Ala        | Val         | Ala<br>860 | His         | Ala         | Lys        | Asp        |
| Va I<br>865 | lle        | Ser        | Asp        | Leu        | Leu<br>870 | Cys        | Asn        | Arg        | He         | Asp<br>875  | l le       | Ser         | Gln         | Leu        | Val<br>880 |
| He          | Thr        | Lys        | Glu        | Leu<br>885 |            | Arg        | Ala        | Ala        | Ala<br>890 |             | Tyr        | Ala         | Gly         | Lys<br>895 | GIn        |
| Ala         | His        | Val        | 900        |            | Ala        | Glu        | Arg        | Met<br>905 |            | ; Lys       | Arg        | Asp         | Pro<br>910  |            | Ser        |
| Ala         | Pro        | Ser<br>915 |            | Gly        | Asp        | Arg        | 920        |            | Tyr        | · Val       | He         | l le<br>925 |             | Ala        | Ala        |
| Lys         | Gly<br>930 |            | Ala        | Ala        | Tyr        | Met<br>935 |            | Ser        | Glu        | Asp         | 940        |             | Phe         | Val        | Leu        |
| Glu<br>945  |            | Ser        | Leu        | ı Pro      | 950        |            | Thr        | · Gin      | ı Tyr      | 955<br>955  |            | Glu         | Gln         | Gln        | Leu<br>960 |
| Ala         | Lys        | : Pro      | Let        | 965        |            | ; lle      | Phe        | e Glu      | 970        |             | Leu        | Gly         | Glu         | 975        | Arg        |
| Ala         | Glu        | ı Ser      | · Val      | Leu        | ı Leu      | ı Are      | Gly        | / Asp      | His        | Thr         | Arg        | Cys         | Lys         | Thr        | Val        |

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#### 109/236

980 985 990

Leu Thr Ser Lys Val Gly Gly Leu Leu Ala Phe Thr Lys Arg Arg Asn 995 1000 1005

Cys Cys IIe Gly Cys Arg Ser Val IIe Asp His Gln Gly Ala Val Cys 1010 1015 1020

Lys Phe Cys Gln Pro Arg Glu Ser Glu Leu Ser Gln Lys Glu Val Ser 1025 1030 1035 1040

His Leu Asn Ala Leu Glu Glu Arg Phe Ser Arg Leu Trp Thr Gln Cys 1045 1050 1055

Gin Arg Cys Gin Gly Ser Leu His Glu Asp Val IIe Cys Thr Ser Arg 1060 1065 1070

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Trp 1105

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Gly Phe Glu Arg Leu Lys Glu Pro Gly Glu Arg Thr Gly Trp Leu IIe 50 55 60

Asn Met His Pro Thr Glu IIe Leu Asp Glu Asp Lys Arg Leu Val Ser 65 70 75 80

Ala Val Asp Tyr Tyr Phe lie Gin Asp Asp Gly Ser Arg Phe Lys Val

Ala Leu Pro Tyr Met Pro Tyr Phe Tyr I le Ala Ala Arg Lys Gly Cys 100 105 110

Asp Arg Glu Val Ser Ser Phe Leu Ser Lys Lys Phe Gln Gly Lys IIe 115 120 125

Ala Lys Leu Glu Asn Val Pro Lys Glu Asp Leu Asp Leu Pro Asn His

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| Glu        | Asp        | Leu        | Val        | Lys<br>165 | Val        | Arg        | Lys        | Glu        | lle<br>170 | Ser        | Pro              | Ala        | Val        | Lys<br>175  | Lys         |
| Asn        | Arg        | Glu        | Gln<br>180 | Asp        | His        | Ala        | Ser        | Asp<br>185 | Glu        | Tyr        | Thr              | Thr        | Met<br>190 | Leu         | Ser         |
| Ser        | He         | Leu<br>195 | Gln        | Gly        | Gly        | Ser        | Val<br>200 | ile        | Thr        | Asp        | Glu              | Glu<br>205 | Glu        | Thr         | Ser         |
| Lys        | Lys<br>210 | He         | Ala        | Asp        | Gln        | Leu<br>215 | Asp        | Asn        | lle        | Val        | Asp<br>220       | Met        | Arg        | Glu         | Tyr         |
| Asp<br>225 | Val        | Pro        | Tyr        | His        | 11e<br>230 | Arg        | Leu        | Ser        | lle        | Asp<br>235 | Leu              | Arg        | lle        | His         | Va I<br>240 |
| Ala        | His        | Trp        | Tyr        | Asn<br>245 | Val        | Arg        | Phe        | Arg        | Gly<br>250 | Asn        | Ala              | Phe        | Pro        | Va I<br>255 | Glu         |
| lle        | Thr        | Arg        | Arg<br>260 | Asp        | Asp        | Leu        | Val        | Glu<br>265 | Arg        | Pro        | Asp              | Pro        | Val<br>270 | Val         | Leu         |
| Ala        | Phe        | Asp<br>275 | He         | Glu        | Thr        | Thr        | Lys<br>280 | Leu        | Pro        | Leu        | Lys              | Phe<br>285 | Pro        | Asp         | Ala         |
| Glu        | Thr<br>290 | Asp        | Gln        | lle        | Met        | Met<br>295 | He         | Ser        | Tyr        | Met        | ile<br>300       | Asp        | Gly        | Gln         | Gly         |
| Tyr<br>305 | Leu        | He         | Thr        | Asn        | Arg<br>310 | Glu        | He         | Val        | Şer        | Glu<br>315 | Asp              | lle        | Glu        | Asp         | Phe<br>320  |
| Glu        | Phe        | Thr        | Pro        | Lys<br>325 | Pro        | Glu        | Tyr        | Glu        | Gly<br>330 | Pro        | Phe              | Cys        | Val        | Phe<br>335  | Asn         |

| Glu        | Pro        | Asp        | Glu<br>340 | Vai                | His        | Leu        | lle        | GIn<br>345 | Arg        | Trp        | Phe         | Glu        | His<br>350 | lle        | Gln        |
|------------|------------|------------|------------|--------------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|
| Glu        | Thr        | Lys<br>355 | Pro        | Thr                | lle        | Met        | Val<br>360 | Thr        | Tyr        | Asn        | Gly         | Asp<br>365 | Phe        | Phe        | Asp        |
| Trp        | Pro<br>370 | Phe        | Val        | Glu                | Ala        | Arg<br>375 | Ala        | Ala        | lle        | His        | Gly<br>380  | Leu        | Ser        | Met        | Tyr        |
| G1n<br>385 | Glu        | He         | Gly        | Phe                | Gln<br>390 | Lys        | Asp        | Ser        | GIn        | Gly<br>395 | Glu         | Tyr        | Lys        | Ala        | Pro<br>400 |
| Gin        | Cys        | lle        | His        | Met<br><b>40</b> 5 | Asp        | Cys        | Leu        | Arg        | Trp<br>410 | Val        | Lys         | Arg        | Asp        | Ser<br>415 | Tyr        |
| Leu        | Pro        |            | Gly<br>420 | Ser                | His        | Asn        | Leu        | Lys<br>425 | Ala        | Ala        | Ala         | Lys        | Ala<br>430 | Lys        | Leu        |
| Gly        | Tyr        | Asp<br>435 | Pro        | Val                | Glu        | Leu        | Asp<br>440 | Pro        | Giu        | Asp        | Met         | Cys<br>445 | Arg        | Met        | Ala        |
| Thr        | Glu<br>450 | Gln        | Pro        | Gin                | Thr        | Leu<br>455 | Ala        | Thr        | Tyr        | Ser        | Va I<br>460 | Ser        | Asp        | Ala        | Val        |
| Ala<br>465 | Thr        | Tyr        | Tyr        | Leu                | Tyr<br>470 | Met        | Lys        | Tyr        | Val        | His<br>475 | Pro         | Phe        | He         | Phe        | Ala<br>480 |
| Leu        | Cys        | Thr        | He         | 11e<br>485         | Pro        | Met        | Glu        | Pro        | Asp<br>490 | Glu        | Val         | Leu        | Arg        | Lys<br>495 | Gly        |
| Ser        | Gly        | Thr        | Leu<br>500 | Cys                | Glu        | Ala        | Leu        | Leu<br>505 | Met        | Val        | GIn         | Ala        | Phe<br>510 | His        | Ala        |
| Asn        | He         | lie<br>515 | Phe        | Pro                | Asn        | Lys        | GIn<br>520 | Glu        | Gln        | Glu        | Phe         | Asn<br>525 | Lys        | Leu        | Thr        |

| Asp        | Asp<br>530 | Gly        | His        | Val        | Leu        | Asp<br>535       | Ala        | Glu        | Thr        | Tyr        | Va I<br>540 | Gly        | Gly         | His        | Val        |
|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|-------------|------------|-------------|------------|------------|
| Glu<br>545 | Ala        | Leu        | Glu        | Ser        | Gly<br>550 | Val              | Phe        | Arg        | Ser        | Asp<br>555 | lle         | Pro        | Cys         | Arg        | Phe<br>560 |
| Arg        | Met        | Asn        | Pro        | Ala<br>565 | Ala        | Phe              | Asp        | Phe        | Leu<br>570 | Leu        | Gin         | Arg        | Val         | Glu<br>575 | Lys        |
| Thr        | Met        | Arg        | His<br>580 | Ala        | He         | Glu              | Glu        | Glu<br>585 | Glu        | Lys        | Val         | Pro        | Va I<br>590 | Glu        | Gin        |
| Ala        | Thr        | Asn<br>595 | Phe        | Gln        | Glu        | Val              | Cys<br>600 | Glu        | Gln        | He         | Lys         | Thr<br>605 | Lys         | Leu        | Thr        |
| Ser        | Leu<br>610 | Lys        | Asp        | Val        | Pro        | Asn<br>615       | Arg        | He         | Glu        | Cys        | Pro<br>620  | Leu        | He          | Tyr        | His        |
| Leu<br>625 | Asp        | Val        | Gly        | Ala        | Met<br>630 | Tyr              | Pro        | Asn        | He         | lle<br>635 | Leu         | Thr        | Asn         | Arg        | Leu<br>640 |
| Gln        | Pro        | Ser        | Ala        | lle<br>645 | Val        | Asp              | Glu        | Ala        | Thr<br>650 | Cys        | Ala         | Ala        | Cys         | Asp<br>655 | Phe        |
| Asn        | Lys        | Pro        | Gly<br>660 | Ala        | Ser        | Cys              | Gln        | Arg<br>665 | Lys        | Met        | Ala         | Trp        | GIn<br>670  | Trp        | Arg        |
| Gly        | Glu        | Phe<br>675 | Met        | Pro        | Ala        | Ser <sup>*</sup> | Arg<br>680 | Ser        | Glu        | Tyr        | His         | Arg<br>685 | He          | GIn        | His        |
| Gln        | Leu<br>690 | Glu        | Ser        | Glu        | Lys        | Phe<br>695       | Pro        | Pro        | Leu        | Phe        | Pro<br>700  | Glu        | Gly         | Pro        | Ala        |
| Arg<br>705 | Ala        | Phe        | His        | Glu        | Leu<br>710 | Ser              | Arg        | Glu        | Glu        | GIn<br>715 | Ala         | Lys        | Tyr         | Glu        | Lys<br>720 |
| Arg        | Arg        | Leu        | Ala        | Asp        | Tyr        | Cys              | Arg        | Lys        | Ala        | Tyr        | Lys         | Lys        | He          | His        | Val        |

|             |            |             |            | 725        |            |            |            |            | 730         |            |            |            |                   | <b>735</b> |                     |
|-------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|-------------------|------------|---------------------|
| Thr         | Lys        | Val         | Glu<br>740 | Glu        | Arg        | Leu        | Thr        | Thr<br>745 | He          | Cys        | Gln        | .Arg       | Glu<br><b>750</b> | Asn        | Ser                 |
| Phe         | Tyr        | Val<br>755  | Asp        | Thr        | Val        | Arg        | Ala<br>760 | Phe        | Arg         | Asp        | Arg        | Arg<br>765 | Tyr               | Glu        | Phe                 |
| Lys         | Gly<br>770 | Leu         | His        | Lys        | Val        | Trp<br>775 | Lys        | Lys        | Lys         | Leu        | Ser<br>780 | Ala        | Ala               | Val        | Glu                 |
| Va I<br>785 | Gly        | Asp         | Ala        | Ser        | Glu<br>790 | Val        | Lys        | Arg        | Cys         | Lys<br>795 | Asn        | Met        | Glu               | He         | Leu<br>8 <b>0</b> 0 |
| Tyr         | Asp        | Ser         | Leu        | GIn<br>805 | Leu        | Ala        | His        | Lys        | Cys<br>810  | lle        | Leu        | Asn        | Ser               | Phe<br>815 | Tyr                 |
| Gly         | Tyr        | Val         | Met<br>820 | Arg        | Lys        | Gly        | Ala        | Arg<br>825 | Trp         | Tyr        | Ser        | Met        | Glu<br>830        | Met        | Ala                 |
| Gly         | ile        | Va I<br>835 | Cys        | Phe        | Thr        | Gly        | Ala<br>840 | Asn        | He          | He         | Thr        | Gin<br>845 | Ala               | Arg        | Glu                 |
| Leu         | lle<br>850 | Glu         | Gln        | He         | Gly        | Arg<br>855 | Pro        | Leu        | Glu         | Leu        | Asp<br>860 | Thr        | Asp               | Gly        | ile                 |
| Trp<br>865  | Cys        | Val         | Leu        | Pro        | Asn<br>870 | Ser        | Phe        | Pro        | Glu         | Asn<br>875 | Phe        | Val        | He                | Lys        | Thr<br>880          |
| Thr         | Asn        | Ala         | Lys        | Lys<br>885 | Pro        | Lys        | Leu        | Thr        | l le<br>890 | Ser        | Tyr        | Pro        | Gly               | Ala<br>895 | Met                 |
| Leu         | Asn        | lle         | Met<br>900 | Val        | Lys        | Glu        | Gly        | Phe<br>905 | Thr         | Asn        | His        | Gln        | Tyr<br>910        | Gin        | Glu                 |
| Leu         | Thr        | Glu         | Pro        | Ser        | Ser        | Leu        | Thr        | Tyr        | Val         | Thr        | His        | Ser        | Glu               | Asn        | Ser                 |

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|                      |             | 915         |             |             |             |                     | 920          |             |             |             |                         | 925         |             |             |             |
|----------------------|-------------|-------------|-------------|-------------|-------------|---------------------|--------------|-------------|-------------|-------------|-------------------------|-------------|-------------|-------------|-------------|
| lle P<br>9           | he<br>30    | Phe         | Glu         | Val         | Asp         | Gly<br>935          | Pro          | Tyr         | Leu         | Ala         | Met <sub>.</sub><br>940 | He          | Leu         | Pro         | Ala         |
| Ser L<br>945         | .ys         | Glu         | Glu         | Gly         | Lys<br>950  | Lys                 | Leu          | Lys         | Lys         | Arg<br>955  | Tyr                     | Ala         | Val         | Phe         | Asn<br>960  |
| Glu A                | lsp         | Gly         | Ser         | Leu<br>965  | Ala         | Glu                 | Leu          | Lys         | Gly<br>970  | Phe         | Glu                     | Val         | Lys         | Arg<br>975  | Arg         |
| Gly 6                | ilu         | Leu         | GIn<br>980  | Leu         | He          | Lys                 | lle          | Phe<br>985  | Gln         | Ser         | Ser                     | Val         | Phe<br>990  | Glu         | Ala         |
| Phe L                | .eu         | Lys<br>995  | Gly         | Ser         | Thr         |                     | Glu<br>1000  | Glu         | Val         | Tyr         |                         | Ser<br>1005 | Val         | Ala         | Lys         |
| Vai <i>I</i>         | Ala<br>010  | Asp         | Tyr         | Trp         |             | <b>A</b> sp<br>1015 | Val          | Leu         | Tyr         |             | Lys<br>1020             | Ala         | Ala         | Asn         | Met         |
| Pro <i>l</i><br>1025 | <b>A</b> sp | Ser         | Glu         |             | Phe<br>1030 | Glu                 | Leu          | lle         |             | Glu<br>1035 | Asn                     | Arg         | Ser         |             | Ser<br>1040 |
| Arg l                | Lys         | Leu         |             | Asp<br>1045 | Tyr         | Gly                 | Glu          |             | Lys<br>1050 | Ser         | Thr                     | Ser         |             | Ser<br>1055 | Thr         |
| Ala l                | Lys         |             | Leu<br>1060 | Ala         | Glu         | Phe                 |              | Gly<br>1065 | Asp         | Gin         | Met                     |             | Lys<br>1070 | Asp         | Ala         |
| Gly I                |             | Ser<br>1075 | Cys         | Arg         | Tyr         |                     | l le<br>1080 | Ser         | Arg         | Lys         |                         | Glu<br>1085 | Gly         | Ser         | Pro         |
| Val <sup>1</sup>     | Thr<br>090  | Glu         | Arg         | Ala         |             | Pro<br>1095         |              | Ala         | lle         |             | Gln<br>1100             |             | Glu         | Pro         | Thr         |
| Val /                |             | Lys         | His         |             | Leu<br>1110 |                     | Lys          | Trp         |             | Lys<br>1115 |                         | Ser         | Ser         |             | GIn<br>1120 |

| •   | Asp                 | He                     | Arg                           | Thr                           | He                       | Leu                      | Asp                    | Trp                           | Asp                           | Tyr                           | Tyr                      | He          | Glu                | Arg                      |
|---|---------------------|------------------------|-------------------------------|-------------------------------|--------------------------|--------------------------|------------------------|-------------------------------|-------------------------------|-------------------------------|--------------------------|-------------|--------------------|--------------------------|
|   |                     | 1                      | 1125                          |                               |                          |                          | •                      | 1130                          |                               |                               |                          | 1           | 135                |                          |
|   |                     |                        |                               |                               |                          |                          |                        |                               |                               |                               |                          |             |                    |                          |
| Leu Gly                                       | Ser                 | Ala                    | He                            | Gln                           | Lys                      | He                       | He                     | Thr                           | He                            | Pro                           | Ala                      | Ala         | Leu                | Gln                      |
|   | •                   | 1140                   |                               |                               |                          |                          | 1145                   |                               |                               |                               | •                        | 1150        |                    |                          |
|   |                     |                        |                               |                               |                          |                          |                        |                               |                               |                               |                          |             |                    |                          |
| Gin Vai                                       | Lys                 | Asn                    | Pro                           | Val                           | Pro                      | Arg                      | Val                    | Lys                           | His                           | Pro                           | Asp                      | Trp         | Leu                | His                      |
|   | 1155                |                        |                               |                               |                          | 1160                     |                        |                               |                               |                               | 1165                     | •           |                    |                          |
|   |                     |                        |                               |                               |                          |                          |                        |                               |                               |                               |                          |             |                    |                          |
| Lys Lys                                       | Leu                 | Leu                    | Glii                          | l vs                          | Aen                      | Asn                      | ما ا                   | Tvr                           | lve                           | Gin                           | l ve                     | Lve         | ما ا               | Sor                      |
| 1170  | Lou                 | Lou                    | u i u                         |                               | 7311<br>1175             | nop                      | 110                    | 1 71                          |                               | 180                           | Lyo                      | Lyo         | 110                | 961                      |
| 1170  |                     |                        |                               |                               | 1175                     |                          |                        |                               | ,                             | 100                           |                          |             |                    |                          |
| Glu Lou                                       | Dho                 | Val                    | Lau                           | Cl.,                          | Glu                      | Lva                      | A                      | e I n                         | Ha                            | Va I                          | Wat                      | Ala 4       | 01 <u>-</u>        | <b>A</b> 1 =             |
| Giu Leu                                       | rne                 | vai                    |                               |                               | uly                      | Lys                      | Arg                    |                               |                               | vai                           | Met                      | AIA         |                    |                          |
| 1185  |                     |                        | ı                             | 190                           |                          |                          |                        | 1                             | 195                           |                               |                          |             | 1                  | 200                      |
|   | _                   | _                      |                               | _                             |                          | _                        | _                      | _                             |                               |                               |                          |             |                    |                          |
| Ser Glu                                       | Asn                 |                        |                               | Ser                           | Leu                      | Cys                      |                        |                               | Asp                           | Met                           | Glu                      |             |                    | Gly                      |
|   |                     | 1                      | 205                           |                               |                          |                          | 1                      | 210                           |                               |                               |                          | 12          | 215                |                          |
|   |                     |                        |                               |                               |                          |                          |                        |                               |                               |                               |                          |             |                    |                          |
| Leu Thr                                       | Lys                 | Pro                    | His                           | His                           | Ser                      | Thr                      | Val                    | Pro                           | Val                           | Ala                           | Thr                      | Lys /       | Arg                | Lys                      |
|   | 1                   | 220                    |                               |                               |                          | 1                        | 225                    |                               |                               |                               | 1                        | 230         |                    |                          |
|   |                     |                        |                               |                               |                          |                          |                        |                               |                               |                               |                          |             |                    |                          |
|   |                     |                        |                               |                               |                          |                          |                        |                               |                               |                               |                          |             |                    |                          |
| Arg Val                                       | Trp                 | Glu                    | Thr                           | Gln                           | Lys                      | Glu                      | Ser                    | Gln                           | Asp                           | lle                           | Ala                      | Leu 1       | Thr '              | Val                      |
|   | Trp<br>1 <b>235</b> | Glu                    | Thr                           | Gln                           |                          | Glu<br>1240              | Ser                    | Gln                           | Asp                           |                               | Ala<br>245               | Leu 1       | Thr '              | Val                      |
|   |                     | Glu                    | Thr                           | Gln                           |                          |                          | Ser                    | Gln                           | Asp                           |                               |                          | Leu 1       | Thr '              | Val                      |
|   | 235                 |                        |                               |                               | 1                        | 1240                     |                        |                               |                               | 1                             | 245                      |             |                    |                          |
| 1   | 235                 |                        |                               | Leu                           | 1                        | 1240                     |                        |                               | Ser                           | 1                             | 245                      |             |                    |                          |
| Pro Trp                                       | 235                 |                        |                               | Leu                           | 1<br>Gly                 | 1240                     |                        |                               | Ser                           | 1<br>Leu                      | 245                      |             |                    |                          |
| Pro Trp<br>1250                               | 1235<br>Gln         | Glu                    | Val                           | Leu<br>1                      | 1<br>Gly<br>1255         | 1240<br>Gln              | Pro                    | Pro                           | Ser<br>1                      | 1<br>Leu<br>260               | 245<br>Gly               | Thr 1       | Thr (              | Gln                      |
| Pro Trp<br>1250<br>Glu Glu                    | 1235<br>Gln         | Glu                    | Val<br>Val                    | Leu<br>1<br>Trp               | 1<br>Gly<br>1255         | 1240<br>Gln              | Pro                    | Pro<br>His                    | Ser<br>1<br>Lys               | 1<br>Leu<br>260               | 245<br>Gly               | Thr 1       | Chr (              | GIn<br>Leu               |
| Pro Trp<br>1250                               | 1235<br>Gln         | Glu                    | Val<br>Val                    | Leu<br>1                      | 1<br>Gly<br>1255         | 1240<br>Gln              | Pro                    | Pro<br>His                    | Ser<br>1                      | 1<br>Leu<br>260               | 245<br>Gly               | Thr 1       | Chr (              | Gln                      |
| Pro Trp<br>1250<br>Glu Glu<br>1265            | GIn<br>Trp          | Glu<br>Leu             | Val<br>Val<br>1               | Leu<br>1<br>Trp<br>270        | 1<br>Gly<br> 255<br>Leu  | Gln<br>Gln               | Pro<br>Phe             | Pro<br>His                    | Ser<br>1<br>Lys<br>275        | 1<br>Leu<br>260<br>Lys        | 245<br>Gly<br>Lys        | Thr 1       | Chr<br>Gin i       | GIn<br>Leu<br>280        |
| Pro Trp<br>1250<br>Glu Glu                    | GIn<br>Trp          | Giu<br>Leu<br>Gin      | Val<br>Val<br>1<br>Arg        | Leu<br>1<br>Trp<br>270        | 1<br>Gly<br> 255<br>Leu  | Gln<br>Gln               | Pro<br>Phe<br>Arg      | Pro<br>His<br>1<br>Lys        | Ser<br>1<br>Lys<br>275        | 1<br>Leu<br>260<br>Lys        | 245<br>Gly<br>Lys        | Thr 1 Trp ( | Chr<br>Gin i<br>1: | GIn<br>Leu<br>280        |
| Pro Trp<br>1250<br>Glu Glu<br>1265            | GIn<br>Trp          | Giu<br>Leu<br>Gin      | Val<br>Val<br>1               | Leu<br>1<br>Trp<br>270        | 1<br>Gly<br> 255<br>Leu  | Gln<br>Gln               | Pro<br>Phe<br>Arg      | Pro<br>His                    | Ser<br>1<br>Lys<br>275        | 1<br>Leu<br>260<br>Lys        | 245<br>Gly<br>Lys        | Thr 1 Trp ( | Chr<br>Gin i       | GIn<br>Leu<br>280        |
| Pro Trp<br>1250<br>Glu Glu<br>1265<br>Gln Ala | GIn<br>Trp<br>GIn   | Glu<br>Leu<br>Gln<br>1 | Val<br>Val<br>1<br>Arg<br>285 | Leu<br>1<br>Trp<br>270<br>Leu | Gly<br>255<br>Leu<br>Ala | Gin<br>Gin<br>Leu        | Pro<br>Phe<br>Arg<br>1 | Pro<br>His<br>1<br>Lys<br>290 | Ser<br>1<br>Lys<br>275<br>Lys | 1<br>Leu<br>260<br>Lys<br>Gln | 245<br>Gly<br>Lys<br>Arg | Thr 1 Trp ( | fhr<br>1:<br>1:    | GIn<br>Leu<br>280<br>Ser |
| Pro Trp<br>1250<br>Glu Glu<br>1265            | GIn Trp GIn         | Glu<br>Leu<br>Gln<br>1 | Val<br>Val<br>1<br>Arg<br>285 | Leu<br>1<br>Trp<br>270<br>Leu | Gly<br>255<br>Leu<br>Ala | Gin<br>Gin<br>Leu<br>Giy | Pro<br>Phe<br>Arg<br>1 | Pro<br>His<br>1<br>Lys<br>290 | Ser<br>1<br>Lys<br>275<br>Lys | 1<br>Leu<br>260<br>Lys<br>Gln | 245<br>Gly<br>Lys<br>Arg | Thr 1 Trp ( | fhr<br>1:<br>1:    | GIn<br>Leu<br>280<br>Ser |

| Gly         |            | Gly<br>1315 | Ser         | Phe         | Leu          |                     | Arg<br>1320 | Thr         | Ala         | Arg         |             | lle<br>1325 | Met         | Asp         | Leu         |
|-------------|------------|-------------|-------------|-------------|--------------|---------------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|
|             | Trp<br>330 | Gln         | lle         | He          |              | l le<br>1335        | Ser         | Glu         | Thr         |             | GIn<br>1340 | Aia         | Gly         | Leu         | Phe         |
| Arg<br>1345 |            | Trp         | Ala         | He          | l le<br>1350 | Gly                 | Asn         | Asp         |             | His<br>1355 | Cys         | ile         | Lys         |             | Ser<br>1360 |
| lle         | Pro        | Arg         |             | Phe<br>1365 | Tyr          | Val                 | Asn         |             | Arg<br>1370 | Val         | Ala         | Lys         |             | Glu<br>1375 | Asp         |
| Gly         | Pro        |             | Tyr<br>1380 | Arg         | Lys          | Val                 |             | Arg<br>1385 | Gly         | Leu         | Phe         |             | Arg<br>1390 | Ser         | Asn         |
| lle         |            | Tyr<br>1395 | Asn         | Leu         | Tyr          |                     | Tyr<br>1400 | Ser         | Val         | Pro         |             | Asp<br>1405 | Met         | Tyr         | Gln         |
|             | His<br>410 | lle         | Asn         | Glu         |              | <b>A</b> sn<br>1415 | Thr         | Glu         | Leu         |             | Val<br>1420 | Pro         | Asp         | He          | Glu         |
| Gly<br>1425 |            | Tyr         | Glu         | Thr         | Gin<br>1430  | Val                 | Pro         | Leu         |             | Phe<br>1435 | Arg         | Ala         | Leu         |             | GIn<br>1440 |
| Leu<br>•    | Gly        | Cys         |             | Cys<br>1445 | Val          | Val                 | Asn         |             | Gin<br>1450 | Leu         | Thr         | Arg         |             | Leu<br> 455 | Ser         |
| Gly         | Trp        |             | Ala<br>1460 | Glu         | Thr          | Phe                 |             | Leu<br>1465 | Glu         | His         | Leu         |             | Met<br>1470 | Arg         | Ser         |
| Leu         |            | Gln<br>1475 | Phe         | Ser         | Tyr          |                     | Glu<br>1480 | Pro         | Gly         | Ser         |             | Arg<br>1485 | His         | He          | Tyr         |
|             | Tyr<br>490 | His         | His         | Thr         |              | Gly<br>1495         | His         | Lys         | Ala         |             | Phe<br>1500 | Gly         | Val         | Phe         | He          |
|             |            |             |             |             |              |                     |             |             |             |             |             |             |             |             |             |

| Pro \$<br>1505 |            | Gin         | Arg         |             | Ala<br>1510 | Ser          | Val                 | Phe         |                     | Leu<br>1515 |             | Thr         | Val         |             | Ser<br>1520  |
|----------------|------------|-------------|-------------|-------------|-------------|--------------|---------------------|-------------|---------------------|-------------|-------------|-------------|-------------|-------------|--------------|
| Asn (          | Gin        | Met         |             | Gly<br>1525 |             | Ser          | Ala                 |             | Tyr<br>1 <b>530</b> |             | Ser         | Glu         |             | Ser<br>1535 | Leu          |
| Leu l          | Leu        |             | Lys<br>1540 |             | Asp         | Pro          |                     | Leu<br>1545 | Leu                 | Pro         | Pro         |             | Lys<br>1550 |             | Thr          |
| Phe (          |            | Val<br>1555 | Arg         | Ala         | Glu         |              | <b>A</b> sn<br>1560 | Leu         | Glu                 | Thr         |             | Cys<br>1565 | Arg         | Ala         | ·lle         |
| Gln /          | Arg<br>570 | Phe         | Leu         | Leu         |             | Tyr<br>1575  | Lys                 | Glu         | Glu                 |             | Arg<br>1580 | Gly         | Pro         | Thr         | Leu          |
| ile /<br>1585  |            | Val         | Gln         |             | Ser<br>1590 | Trp          | Glu                 | Leu         |                     | Arg<br>1595 | Leu         | Thr         | Ser         |             | l le<br>1600 |
| Pro \          | Vai        | Leu         |             | Glu<br>1605 | Phe         | Pro          | Leu                 |             | Pro<br>1610         | lle         | Arg         | Val         |             | Asp<br>1615 | Lys          |
| lle S          | Ser        |             | Ala<br>1620 | Val         | Leu         | Asp          |                     | Gln<br>1625 | Arg                 | His         | Gly         |             | Arg<br>1630 |             | Met          |
| lle #          |            | His<br>1635 | Tyr         | Leu         | Asn         |              | Asp<br>1640         | Leu         | Cys                 | Leu         |             | Gln<br>1645 | Ala         | Phe         | Glu          |
| Met S          | Ser<br>650 | Arg         | Tyr         | Phe         |             | l le<br>1655 | Pro                 | Val         | Gly                 |             | Leu<br>1660 | Pro         | Glu         | Asp         | He           |
| Ser 1<br>1665  | lle        | Phe         | Gly         |             | Asp<br>1670 | Leu          | Phe                 | Phe         |                     | Arg<br>1675 | His         | Leu         | Gln         |             | His<br>1680  |
| Asn 1          | lis        | Leu         |             | Trp<br>1685 | Leu         | Ser          | Pro                 |             | Ser<br>1690         | Arg         | Pro         | Asp         |             | Gly<br>1695 | Giy          |
| lvs 6          | alis       | Ala         | Aen         | Asn         | Asn         | Arø          | Leu                 | Val         | Met                 | Glo         | Phe         | Asn         | Asp         | Arø         | Ala          |

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|                 | 1710        | •           |             |             |             | 1705                 | -           |                     |             |             | 700         |             |               |           |
|-----------------|-------------|-------------|-------------|-------------|-------------|----------------------|-------------|---------------------|-------------|-------------|-------------|-------------|---------------|-----------|
| Val Glu         | Cys         | Vai<br>1725 | •           | Ser         | Tyr         | Cys                  | Giy<br>1720 |                     | Ser         | Asn         | ile         | Glu<br>1715 |               | Thi       |
| His His         | Ser         | Gln         | Leu<br>1740 |             | Thr         | Asn                  | Val         | Ala<br>1735         |             | Asn         | Gln         | lle         | . Asp<br>1730 | Lei       |
| Asp Val<br>1760 | Phe         | Ser         | He          | Gly<br>1755 |             | Ser                  | Gly         | Ala                 | Gly<br>1750 |             | Met         | Asp         |               | Va<br>174 |
| Ala Ala<br>775  |             | Asn         | Gly         | Thr         | Val<br>1770 |                      | Asp         | Glu                 | Leu         | Ser<br>1765 |             | Gln         | e GIn         | П         |
| Thr Phe         | Ser<br>1790 |             | Cys         | Leu         | Ala         | Thr<br>1 <b>7</b> 85 |             | Asp                 | Tyr         | Asn         | Ala<br>1780 |             | ^ Ala         | Sei       |
| Thr Gin         | He          | Glu<br>1805 |             | Val         | Trp         | Gly                  | Va)<br>1800 |                     | Met         | Ser         | Lys         | Leu<br>1795 |               | Arı       |
| Arg Trp         | Tyr         | Phe         | His<br>1820 |             | Val         | Gln                  | Asn         | <b>A</b> sp<br>1815 |             | Tyr         | lle         | Asn         | - His<br>1810 | Tyı       |
| Arg Thr<br>1840 | His         | Leu         | Ala         | Pro<br>1835 |             | His                  | Leu         | Leu                 | Ser<br>1830 |             | Pro         | Ser         |               | Le:       |
| Glu Phe<br>855  |             | lle         | Leu         | Gln         | Leu<br>1850 |                      | Leu         | Lys                 | Lys         | Met<br>1845 |             | Asn         | aiH u         | Le        |
| lle ile         | Arg<br>1870 |             | Phe         | Asn         | Ala         | Tyr<br>1865          |             | Val                 | Ser         | Ser         | Gly<br>1860 |             | s Arg         | Ly        |
| <b>V</b> al Glu | Tyr         | Ala<br>1885 |             | Ala         | Asp         | Glu                  | lle<br>1880 |                     | Arg         | Lys         | Lys         | Thr<br>1875 |               | Le        |
| Leu Thr         | Ser         | His         | Phe<br>1900 |             | Glu         | Lys                  | Ser         | His<br>1895         |             | Ser         | Asn         | Thr         | - Ile<br>1890 | Ty        |

| lle Ser<br>1905 | Phe Ser         |               | Cys<br>910  | lrp         | Glu         | Pne         |             | Leu<br> 915 | Irp         | Met         | Asp         |             | Ser<br>920  |
|-----------------|-----------------|---------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|
| Asn Tyr         | Gly Gly         | lle  <br>1925 | Lys (       | Gly         | Lys         |             | Pro<br>1930 | Ser         | Ser         | He          |             | Cys<br>1935 | Gly         |
| Gin Vai         | Lys Glu<br>1940 |               | Asp :       | Ser         |             | Ala<br>945  | Arg         | Glu         | Glu         |             | Asp<br>1950 | Glu         | Glu         |
| Glu Glu<br>. 1  | Asp Lys<br>955  | Glu           | Lys /       |             | Glu<br>960  | Glu         | Glu         | Glu         |             | Met<br>1965 | Gly         | Glu         | Ser         |
| Glu Val<br>1970 | Glu Ası         | Leu           |             | Glu<br>975  | Asn         | Asn         | Trp         |             | lle<br>1980 | Leu         | Gln         | Phe         | Leu         |
| Pro GIn<br>1985 | Ala Ala         |               | Cys<br>990  | Gin         | Ser         | Tyr         |             | Leu<br>1995 | Met         | He          | Val         |             | Ala<br>2000 |
| Tyr lle         | Val Ala         | a Val<br>2005 | Tyr         | Gln         | Ser         |             | Lys<br>2010 | Glu         | Glu         | Leu         |             | His<br>2015 | Ser         |
| Ala Pro         | Gly Sei<br>2020 |               | Pro         | Val         |             | Arg<br>2025 | Lys         | Gly         | Ala         |             | GIn<br>2030 | Phe         | Ser         |
| Gln Glu         | Ser Gli<br>2035 | ı Gly         | Ala         |             | Gly<br>2040 | Ser         | Leu         | Pro         |             | Met<br>2045 | lle         | Thr         | Phe         |
| Ser Gin<br>2050 | Asp Ty          | - Val         |             | Asn<br>:055 | Glu         | Leu         | Thr         |             | Ser<br>2060 | Phe         | Phe         | Thr         | He          |
| Thr Gln<br>2065 | Lys II          |               | Lys<br>2070 | Lys         | Val         | Thr         |             | Ser<br>2075 | Arg         | Asn         | Thr         |             | Glu<br>2080 |
|                 |                 |               |             |             |             |             |             |             |             |             |             |             |             |

|                     | Leu Glu Phe<br>2100 | lle Lys Tyr<br>2105 | Val Cys Lys         | Val Leu Ser Glr<br>2110 |
|---------------------|---------------------|---------------------|---------------------|-------------------------|
| Asp Thr Asn<br>2115 | lle Thr Asn         | Gin Vai Asn<br>2120 |                     | Arg Asp Leu Leu<br>2125 |
| Arg Leu Val<br>2130 |                     | Glu Phe Ser<br>2135 | Glu Glu Ala<br>2140 | Gin Phe Arg Asp         |
| Pro Cys His<br>2145 | Ser Tyr Val<br>2150 | Leu Pro Glu         | Val lie Cys<br>2155 | His Ser Cys Asn<br>2160 |
| Phe Cys Arg         | Asp Leu Asp<br>2165 |                     | Asp Ser Ser<br>2170 | Phe Ser Gln Asp<br>2175 |
|                     | Leu Pro Gln<br>2180 | Trp Leu Cys<br>2185 | Ser Asn Cys         | Gin Ala Pro Tyr<br>2190 |
| Asp Ser Ser<br>2195 | Ala ile Glu         | Ser Ala Leu<br>2200 |                     | Leu Gin Arg Lys<br>2205 |
| Leu Met Ala<br>2210 | •                   | Gln Asp Leu<br>2215 | Val Cys Leu<br>2220 | Lys Cys Arg Gly         |
| Met Lys Glu<br>2225 | Thr His Met<br>2230 | Pro Val Tyr         | Cys Ser Cys<br>2235 | Ala Gly Asp Phe<br>2240 |
| Thr Leu Thr         | lie Arg Thr<br>2245 |                     | Met Glu Gin<br>2250 | lle Arg lle Phe<br>2255 |
|                     | Ala Lys Tyr<br>2260 | Tyr Ser Met<br>2265 | Ser Tyr Leu         | Gin Giu Thr ile<br>2270 |
| Glu Trp Leu<br>2275 | Leu Gin Thr         | Ser Pro Val         | Ser Asn Cys         |                         |

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<211> 3325

<212> DNA

<213> Rattus norvegicus

#### <400> 59

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<210> 60 <211> 1103 <212> PRT

<213> Rattus norvegicus

<400> 60

Met Asp Gly Lys Arg Arg Gln Ala Pro Ser Ser Gly Val Pro Pro Lys 1 5 10 15

Arg Ala Cys Lys Gly Leu Trp Asp Glu Asp Glu Pro Ser Gln Phe Glu

|            |            |            | 20         |            |            |            |            | 25         |            |            |            |            | 30         |            |            |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| Glu        | Asn        | Leu<br>35  | Ala        | Leu        | Leu        | Glu        | Glu<br>40  | He         | Glu        | Ala        | Glu        | Asn<br>45  | Arg        | Leu        | Gin        |
| Glu        | Ala<br>50  | Glu        | Glu        | Glu        | Leu        | GIn<br>55  | Leu        | Pro        | Pro        | Glu        | Gly<br>60  | lle        | Val        | Gly        | Gly        |
| GIn<br>65  | Phe        | Ser        | Thr        | Ala        | Asp<br>70  | He         | Asp        | Pro        | Arg        | Trp<br>75  | Leu        | Arg        | Pro        | Thr        | Pro<br>80  |
| Leu        | Ala        | Leu        | Asp        | Pro<br>85  | Ser        | Thr        | Glu        | Pro        | Leu<br>90  | lle        | Phe        | Gin        | Gln        | Leu<br>95  | Glu        |
| He         | Asp        | His        | Tyr<br>100 | Val        | Gly        | Thr        | Ser        | Pro<br>105 | Pro        | Leu        | Pro        | Glu        | Gly<br>110 | Pro        | Pro        |
| Ala        | Ser        | Arg<br>115 | Asn        | Ser        | Val        | Pro        | lle<br>120 | Leu        | Arg        | Ala        | Phe        | Gly<br>125 | Val        | Thr        | Asp        |
| Glu        | Gly<br>130 | Phe        | Ser        | Val        | Cys        | Cys<br>135 | His        | lle        | His        | Gly        | Phe<br>140 | Ala        | Pro        | Tyr        | Phe        |
| Tyr<br>145 | Thr        | Pro        | Ala        | Pro        | Pro<br>150 | Gly        | Phe        | Gly        | Ala        | Glu<br>155 | His        | Leu        | Ser        | Glu        | Leu<br>160 |
| Gin        | Arg        | Glu        | Leu        | Asn<br>165 | Ala        | Ala        | He         | Ser        | Arg<br>170 | Asp        | GIn        | Arg        | Gly        | Gly<br>175 | Lys        |
| Glu        | Leu        | Ser        | Gly<br>180 | Pro        | Ala        | Val        | Leu        | Ala<br>185 | He         | Glu        | Leu        | Cys        | Ser<br>190 | Arg        | Glu        |
| Ser        | Met        | Phe<br>195 | Gly        | Tyr        | His        | Gly        | His<br>200 | Gly        | Pro        | Ser        | Pro        | Phe<br>205 | Leu        | Arg        | He         |
| Thr        | Leu<br>210 | Ala        | Leu        | Pro        | Arg        | Leu<br>215 | Met        | Ala        | Pro        | Ala        | Arg<br>220 | Arg        | Leu        | Leu        | Glu        |

| GIn<br>225 | Gly        | lle        | Arg        | Val        | Pro<br>230 | Gly        | Leu        | Gly        | Thr         | Pro<br>235 | Ser        | Phe        | Ala         | Pro        | Tyr<br>240 |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|
| Glu        | Ala        | Asn        | Val        | Asp<br>245 | Phe        | Glu        | lle        | Arg        | Phe<br>250  | Met        | Val        | Asp        | Ala         | Asp<br>255 | ile        |
| Val        | Gly        | Cys        | Asn<br>260 | Trp        | Leu        | Glu        | Leu        | Pro<br>265 | Ala         | Gly        | Lys        | Tyr        | Va I<br>270 | Arg        | Arg        |
| Ala        | Glu        | Lys<br>275 | Lys        | Ala        | Thr        | Leu        | Cys<br>280 | Gin        | Leu         | Glu        | Val        | Asp<br>285 | Val         | Leu        | Trp        |
| Ser        | Asp<br>290 | Val        | ile        | Ser        | His        | Pro<br>295 | Pro        | Glu        | Gly         | Gin        | Trp<br>300 | Gin        | Arg         | He         | Ala        |
| Pro<br>305 | Leu        | Arg        | Val        | Leu        | Ser<br>310 | Phe        | Asp        | lle        | Glu         | Cys<br>315 | Ala        | Gly        | Arg         | Lys        | Gly<br>320 |
| lle        | Phe        | Pro        | Glu        | Pro<br>325 | Glu        | Arg        | Asp        | Pro        | Va I<br>330 | lle        | Gin        | He         | Cys         | Ser<br>335 | Leu        |
| Gly        | Leu        | Arg        | Trp<br>340 | Gly        | Glu        | Pro        | Glu        | Pro<br>345 | Phe         | Leu        | Arg        | Leu        | Ala<br>350  | Leu        | Thr        |
| Leu        | Arg        | Pro<br>355 | Cys        | Ala        | Pro        | He         | Leu<br>360 | Gly        | Ala         | Lys        | Val        | GIn<br>365 | Ser         | Tyr        | Glu        |
| Arg        | Glu<br>370 | Glu        | Asp        | Leu        | Leu        | GIn<br>375 | Ala        | Trp        | Ala         | Thr        | Phe<br>380 | He         | Leu         | Ala        | Met        |
| Äsp<br>385 | Pro        | Asp        | Val        | He         | Thr<br>390 | Gly        | Tyr        | Asn        | He          | GIn<br>395 | Asn        | Phe        | Asp         | Leu        | Pro<br>400 |
| Tyr        | Leu        | lle        | Ser        | Arg<br>405 | Aia        | Gln        | Thr        |            | Lys<br>410  | Val        | Asp        | Arg        |             | Pro<br>415 | Phe        |

| Leu        | Gly        | Arg        | Va I<br>420 | Thr        | Gly        | Leu        | Arg                | Ser<br>425 | Asn        | lle        | Arg        | Asp         | Ser<br>430 | Ser        | Phe        |
|------------|------------|------------|-------------|------------|------------|------------|--------------------|------------|------------|------------|------------|-------------|------------|------------|------------|
| Gln        | Ser        | Arg<br>435 | Gin         | Val        | Gly        | Arg        | Arg<br>440         | Asp        | Ser        | Lys        | Va I       | Va I<br>445 | Ser        | Met        | Val        |
| Gly        | Arg<br>450 | Val        | Gln         | Met        | Asp        | Met<br>455 | Leu                | GIn        | Val        | Leu        | Leu<br>460 | Arg         | Glu        | Tyr        | Lys        |
| Leu<br>465 | Arg        | Ser        | Tyr         | Thr        | Leu<br>470 | Asn        | Ala                | Val        | Ser        | Phe<br>475 | His        | Phe         | Leu        | Gly        | Glu<br>480 |
| Gln        | Lys        | Glu        | Asp         | Val<br>485 | GIn        | His        | Ser                | lle        | lle<br>490 | Thr        | Asp        | Leu         | Gln        | Asn<br>495 | Gly        |
| Asn        | Glu        | Gin        | Thr<br>500  | Arg        | Arg        | Arg        | Leu                | A1a<br>505 | Val        | Tyr        | Cys        | Leu         | Lys<br>510 | Asp        | Ala        |
| Phe        | Leu        | Pro<br>515 | Leu         | Arg        | Leu        | Leu        | Glu<br><b>52</b> 0 | Arg        | Leu        | Met        | Val        | Leu<br>525  | Val        | Asn        | Asn        |
| Vai        | Glu<br>530 | Met        | Ala         | Arg        | Val        | Thr<br>535 | Gly                | Val        | Pro        | Leu        | Gly<br>540 | Tyr         | Leu        | Leu        | Ser        |
| Arg<br>545 | Gly        | Gin        | Gln         | Val        | Lys<br>550 | Val        | Val                | Ser        | Gin        | Leu<br>555 | Leu        | Arg         | GIn        | Ala        | Met<br>560 |
| Arg        | Glu        | Gly        | Leu         | Leu<br>565 | Met        | Pro        | Val                | Val        | Lys<br>570 | Thr        | Glu        | Gly         | Gly        | Glu<br>575 | Asp        |
| Tyr        | Thr        | Gly        | Ala<br>580  | Thr        | Vai        | lle        | Glu                | Pro<br>585 | Leu        | Lys        | Gly        | Tyr         | Tyr<br>590 | Asp        | Val        |
| Pro        | lle        | Ala<br>595 | Thr         | Leu        | Asp        | Phe        | Ser<br>600         | Ser        | Leu        | Tyr        | Pro        | Ser<br>605  | He         | Met        | Met        |
| Ala        | His        | Asn        | Leu         | Cys        | Tyr        | Thr        | Thr                | Leu        | Leu        | Arg        | Pro        | Gly         | Ala        | Ala        | Gln        |

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|            | 610        |             |            |            |            | 615        |            |            |            |            | 620              |            |            |            |                    |
|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|--------------------|
| Lys<br>625 | Leu        | Gly         | Leu        | Lys        | Pro<br>630 | Asp        | Glu        | Phe        | lle        | Lys<br>635 | Thr <sub>.</sub> | Pro        | Thr        | Gly        | <b>A</b> sp<br>640 |
| Glu        | Phe        | Val         | Lys        | Ala<br>645 | Ser        | Vai        | Arg        | Lys        | Gly<br>650 | Leu        | Leu              | Pro        | GIn        | lle<br>655 | Leu                |
| Glu        | Asņ        | Leu         | Leu<br>660 | Ser        | Ala        | Arg        | Lys        | Arg<br>665 | Ala        | Lys        | Ala              | Glu        | Leu<br>670 | AÌa        | GIn                |
| Glu        | Thr        | Asp<br>675  | Pro        | Leu        | Arg        | Arg        | GIn<br>680 | Val        | Leu        | Asp        | Gly              | Arg<br>685 | Gln        | Leu        | Ala                |
| Leu        | Lys<br>690 | Val         | Ser        | Pro        | Asn        | Ser<br>695 | Val        | Tyr        | Gly        | Phe        | Thr<br>700       | Gly        | Ala        | GIn        | Val                |
| Gly<br>705 | Lys        | Leu         | Pro        | Cys        | Leu<br>710 | Glu        | lle        | Ser        | Gin        | Ser<br>715 | Val              | Thr        | Gly        | Phe        | Gly<br>720         |
| Arg        | GIn        | Met         | lle        | Glu<br>725 | Lys        | Thr        | Lys        | Gin        | Leu<br>730 | Val        | Glu              | Thr        | Lys        | Tyr<br>735 | Thr                |
| Leu        | Glu        | Asn         | Gly<br>740 | Tyr        | Asp        | Ala        | Asn        | Ala<br>745 | Lys        | Val        | Val              | Tyr        | Gly<br>750 | Asp        | Thr                |
| Asp        | Ser        | Va I<br>755 | Met        | Cys        | Arg        | Phe        | Gly<br>760 | Val        | Ser        | Ser        | Val              | Ala<br>765 | Glu        | Ala        | Met                |
| Ser        | Leu<br>770 | Gly         | Arg        | Glu        | Ala        | Ala<br>775 | Asn        | Trp        | Vai        | Ser        | Ser<br>780       | His        | Phe        | Pro        | Ser                |
| Pro<br>785 | He         | Arg         | Leu        | Glu        | Phe<br>790 | Glu        | Lys        | Val        | Tyr        | Phe<br>795 | Pro              | Tyr        | Leu        | Leu        | l le<br>800        |
| Ser        | Lys        | Lys         | Arg        | Tyr<br>805 | Ala        | Gly        | Leu        | Leu        | Phe<br>810 | Ser        | Ser              | Arg        | Ser        | Asp<br>815 | Ala                |

| His        | Asp        | Arg         | Met<br>820 | Asp         | Cys                | Lys        | Gly         | Leu<br>825 | Glu        | Ala                 | Val        | Arg         | Arg<br>830  | Asp        | Asn        |
|------------|------------|-------------|------------|-------------|--------------------|------------|-------------|------------|------------|---------------------|------------|-------------|-------------|------------|------------|
| Cys        | Pro        | Leu<br>835  | Val        | Ala         | Asn                | Leu        | Va I<br>840 | Thr        | Ser        | Ser                 | Leu        | Arg<br>845  | Arg         | He         | Leu        |
| Val        | Asp<br>850 | Arg         | Asp        | Pro         | Asp                | Gly<br>855 | Ala         | Val        | Ala        | His                 | Ala<br>860 | Lys         | Asp         | Val        | lle        |
| Ser<br>865 | Asp        | Leu         | Leu        | Cys         | <b>A</b> sn<br>870 | Arg        | lle         | Asp        | He         | Ser<br>875          | Gln        | Leu         | Val         | lle        | Thr<br>880 |
| Lys        | Glu        | Leu         | Thr        | Arg<br>885  | Ala                | Ala        | Ala         | Asp        | Tyr<br>890 | Ala                 | Gly        | Lys         | Gin         | Ala<br>895 | His        |
| Val        | Glu        | Leu         | Ala<br>900 | Glu         | Arg                | Met        | Arg         | Lys<br>905 | Arg        | Asp                 | Pro        | Gly         | Ser<br>910  | Ala        | Pro        |
| Asn        | Leu        | Gly<br>915  | Asp        | Arg         | Val                | Pro        | Tyr<br>920  | Val        | He         | lle                 | Gly        | Ala<br>925  | Ala         | Lys        | Gly        |
| Val        | Ala<br>930 | Ala         | Tyr        | Met         | Lys                | Ser<br>935 | Glu         | Asp        | Pro        | Leu                 | Phe<br>940 | Val         | Leu         | Glu        | His        |
| Ser<br>945 | Leu        | Pro         | lle        | Asp         | Thr<br>950         | GIn        | Tyr         | Tyr        | Leu        | Glu<br>9 <b>5</b> 5 | Gin        | Gin         | Leu         | Ala        | Lys<br>960 |
| Pro        | Leu        | Leu         | Arg        | l le<br>965 | Phe                | Glu        | Pro         | lle        | Leu<br>970 | Gly                 | Glu        | Gly         | Arg         | Ala<br>975 | Glu        |
| Ser        | Val        | Leu         | Leu<br>980 | Arg         | Gly                | Asp        | His         | Thr<br>985 | Arg        | Cys                 | Lys        | Thr         | Va I<br>990 | Leu        | Thr        |
| Ser        | Lys        | Va I<br>995 | Gly        | Gly         | Leu                |            | Ala<br>1000 | Phe        | Thr        | Lys                 |            | Arg<br>1005 | Asn         | Ser        | Cys        |

lle Gly Cys Arg Ser Val IIe Asp His Gln Gly Ala Val Cys Lys Phe 1010 1015 1020

Cys Gln Pro Arg Glu Ser Glu Leu Tyr Gln Lys Glu Val Ser His Leu 1025 1030 1035 1040

Asn Ala Leu Glu Giu Arg Phe Ser Arg Leu Trp Thr Gin Cys Gln Arg 1045 1050 1055

Cys Gln Gly Ser Leu His Glu Asp Val 11e Cys Thr Ser Arg Asp Cys 1060 1065 1070

Pro IIe Phe Tyr Met Arg Lys Lys Val Arg Lys Asp Leu Glu Asp Gln 1075 1080 1085

Glu Arg Leu Leu Gln Arg Phe Gly Pro Pro Gly Pro Glu Ala Trp 1090 1095 1100

<210> 61

<211> 3451

<212> DNA

<213> Bos taurus

#### **<400> 61**

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| acggcccctc | cccgtttctg | cgtatcacct | tggcactgcc | ccgcctcatg | gcacctgccc | 720  |
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| acgaggccaa | cgttgacttt | gagatccggt | tcatggtgga | cacggacatc | gtgggctgca | 840  |
| actggctgga | gctcccagcc | gggaaataca | tcctgaggcc | ggaggggaag | gccactctgt | 900  |
| gtcagctgga | ggccgacgtg | ctgtggtcag | acgtgatcag | ccacccgccg | gaaggagagt | 960  |
| ggcagcgaat | cgcccctctg | cgcgtgctca | gcttcgacat | cgagtgcgct | ggtcgcaaag | 1020 |
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| ggggcgagcc | ggagcccttc | ctgcgcctgg | cgctcaccct | gcggccctgc | gcccccatcc | 1140 |
| tgggcgccaa | ggtgcagagc | tatgagcggg | aggaggacct | gctccaggcc | tggtcgacct | 1200 |
| tcatccgcat | catggatccc | gatgtgatca | ccggctacaa | tatccagaac | tttgaccttc | 1260 |
| cctacctcat | ctcccgggcc | cagaccctca | aggtgccagg | cttccccttg | ctgggccgtg | 1320 |
| tgattggcct | ccgctccaac | atccgggagt | cgtccttcca | gtccaggcag | actggccggc | 1380 |
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| agcagaagga | ggacgtgcag | cacagcatca | tcacagacct | gcagaacggt | aacgaccaga | 1560 |
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| ccacggtcat | cgagccgctg | aaagggtact | acgacgttcc | catcgccacc | ttggacttct | 1860 |
| cctcgctgta | cccgtccatc | atgatggccc | acaacctgtg | ctacaccaca | ctcctgcggc | 1920 |
| ccggggccgc | ccagaaactg | ggcctgaccg | aggatcagtt | catcaagacg | cccacggggg | 1980 |
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| tcagcgcccg | gaagagggcc | aaggccgagc | tggccaagga | gacagacccc | ctacggcggc | 2100 |
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| ctggcgccca | ggtgggcagg | ctcccgtgcc | tggaaatctc | acagagtgtc | accgggttcg | 2220 |
| ggcgccagat | gattgagaag | acaaagcagc | ttgtggagac | caagtacacg | gtggaaaacg | 2280 |
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| tcagcaagaa | gcgttacgca | ggcctgctct | tctcctcccg | gccggacgcc | cacgaccgca | 2520 |
| tggactgcaa | gggcctggag | gccgtgcgca | gggacaactg | cccctggtg  | gccaacctcg | 2580 |
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| tgcgcgggga | ccacactcgc | tgcaagacgg | tgctcacggg | gaaggtgggc | ggcctcctgg | 3060 |
| ccttcgccaa | acgccggaac | tgctgcatcg | gctgccgcac | tgtcctcagc | caccagggag | 3120 |
| ccgtgtgcaa | gttctgccag | ccccgggagt | cagagctgta | ccagaaggag | gtgtcccacc | 3180 |
| tgagtgccct | ggaggagcga | ttctcacgcc | tgtggacgca | gtgccagcgc | tgccagggca | 3240 |
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| cagaggcttg | gtgacctctg | acctcaacga | acttcccacc | ttgggggcgc | gggggggaca | 3420 |
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<211> 1106

<212> PRT

<213> Bos taurus

<400> 62

Met Asp Gly Lys Arg Arg Pro Gly Pro Gly Pro Gly Val Pro Pro Lys

1 5 10 15

Arg Ala Arg Gly Gly Leu Trp Asp Glu Asp Glu Ala Tyr Arg Pro Ser 20 25 30

Gin Phe Giu Giu Leu Ala Leu Met Giu Giu Met Giu Ala Giu Arg 35 40 45

Arg Leu Gin Giu Giu Giu Giu Giu Leu Gin Ser Ala Leu Giu Ala 50 55 60

Ala Asp Gly Gln Phe Ser Pro Thr Ala IIe Asp Ala Arg Trp Leu Arg 65 70 75 80

Pro Ala Pro Pro Ala Leu Asp Pro Gin Met Glu Pro Leu IIe Phe Gin 85 90 95

Gin Leu Glu ile Asp His Tyr Val Ala Pro Ala Arg Pro Leu Pro Gly

PCT/JP2004/004378

|            |            |            | 100        |            |            |             |            | 105        |            |            |             |            | 110        |             |            |
|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|-------------|------------|------------|-------------|------------|
| Ala        | Pro        | Pro<br>115 | Pro        | Ser        | Gln        | Asp         | Ser<br>120 | Val        | Pro        | He         | Leu         | Arg<br>125 | Ala        | Phe         | Gly        |
| Val        | Thr<br>130 | Asn        | Glu        | Gly        | Val        | Ser<br>135  | Vai        | Cys        | Cys        | His        | l le<br>140 | His        | Gly        | Phe         | Ala        |
| Pro<br>145 | Tyr        | Phe        | Ţyr        | Thr        | Pro<br>150 | Ala         | Pro        | Pro        | Gly        | Phe<br>155 | Gly         | Pro        | Glu        | His         | Leu<br>160 |
| Ser        | Glu        | Leu        | GIn        | Arg<br>165 | Glu        | Leu         | Ser        | Ala        | Ala<br>170 | He         | Ser         | Arg        | Asp        | GIn<br>175  | Arg        |
| Gly        | Gly        | Lys        | Glu<br>180 | Leu        | Thr        | Gly         | Pro        | Ala<br>185 | Val        | Leu        | Ala         | Val        | Glu<br>190 | Leu         | Cys        |
| Ser        | Arg        | Glu<br>195 | Ser        | Met        | Phe        | Gly         | Tyr<br>200 | His        | Gly        | His        | Gly         | Pro<br>205 | Ser        | Pro         | Phe        |
| Leu        | Arg<br>210 | He         | Thr        | Leu        | Ala        | Leu<br>215  | Pro        | Arg        | Leu        | Met        | Ala<br>220  | Pro        | Ala        | Arg         | Arg        |
| Leu<br>225 | Leu        | Glu        | GIn        | Gly        | lle<br>230 | Arg         | Leu        | Ala        | Gly        | Leu<br>235 | Gly         | Thr        | Pro        | Ser         | Phe<br>240 |
| Ala        | Pro        | Tyr        | Glu        | Ala<br>245 | Asn        | Val         | Asp        | Phe        | Glu<br>250 | He         | Arg         | Phe        | Met        | Va I<br>255 | Asp        |
| Thr        | Asp        | He         | Va!<br>260 | Gly        | Cys        | Asn         | Trp        | Leu<br>265 | Glu        | Leu        | Pro         | Ala        | Gly<br>270 | Lys         | Tyr        |
| He         | Leu        | Arg<br>275 | Pro        | Glu        | Gly        | Lys         | Ala<br>280 | Thr        | Leu        | Cys        | Gln         | Leu<br>285 | Glu        | Ala         | Asp        |
| Val        | Leu<br>290 | Trp        | Ser        | Asp        | Val        | l ie<br>295 | Ser        | His        | Pro        | Pro        | Glu<br>300  | Gly        | Glu        | Trp         | Gln        |

| Arg<br>305 | He         | Ala        | Pro        | Leu        | Arg<br>310 | Val        | Leu        | Ser        | Phe        | Asp<br>315 | lle        | Glu        | Cys        | Ala        | Gly<br>320 |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| Arg        | Lys        | Gly        | He         | Phe<br>325 | Pro        | Glu        | Pro        | Glu        | Arg<br>330 | Asp        | Pro        | Val        | He         | GIn<br>335 | He         |
| Cys        | Ser        | Leu        | Gly<br>340 | Leu        | Arg        | Trp        | Gly        | Glu<br>345 | Pro        | Glu        | Pro        | Phe        | Leu<br>350 | Arg        | Leu        |
| Ala        | Leu        | Thr<br>355 | Leu        | Arg        | Pro        | Cys        | Ala<br>360 | Pro        | He         | Leu        | Gly        | Ala<br>365 | Lys        | Val        | Gln        |
| Ser        | Tyr<br>370 | Glu        | Arg        | Glu        | Glu        | Asp<br>375 | Leu        | Leu        | Gin        | Ala        | Trp<br>380 | Ser        | Thr        | Phe        | lle        |
| Arg<br>385 | lle        | Met        | Asp        | Pro        | Asp<br>390 | Val        | He         | Thr        | Gly        | Tyr<br>395 | Asn        | He         | Gin        | Asn        | Phe<br>400 |
| Asp        | Leu        | Pro        | Tyr        | Leu<br>405 | He         | Ser        | Arg        | Ala        | GIn<br>410 | Thr        | Leu        | Lys        | Val        | Pro<br>415 | Gly        |
| Phe        | Pro        | Leu        | Leu<br>420 | Gly        | Arg        | Val        | He         | Gly<br>425 | Leu        | Arg        | Ser        | Asn        | lle<br>430 | Arg        | Glu        |
| Şer        | Ser        | Phe<br>435 | GIn        | Ser        | Arg        | Gin        | Thr<br>440 | Gly        | Arg        | Arg        | Asp        | Ser<br>445 | Lys        | Val        | Val        |
| Ser        | Met<br>450 | Val        | Gly        | Arg        | Val        | GIn<br>455 | Met        | Asp        | Met        | Leu        | GIn<br>460 | Val        | Leu        | Leu        | Arg        |
| Glu<br>465 | Tyr        | Lys        | Leu        | Arg        | Ser<br>470 | Tyr        | Thr        | Leu        | Asn        | Ala<br>475 | Val        | Ser        | Phe        | His        | Phe<br>480 |
| Leu        | Gly        | Glu        | Gln        | Lys<br>485 | Glu        | Asp        | Val        | GIn        | His<br>490 | Ser        | lle        | lle        | Thr        | Asp<br>495 | Leu        |

| Gin        | Asn        | Gly         | Asn<br>500 | Asp        | Gin        | Thr        | Arg        | Arg<br>505 | Arg         | Leu         | Ala         | Vai        | Tyr<br>510 | Cys        | Leu        |
|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|-------------|-------------|------------|------------|------------|------------|
| Lys        | Asp        | Ala<br>515  | Phe        | Leu        | Pro        | Leu        | Arg<br>520 | Leu        | Leu         | Glu         | Arg         | Leu<br>525 | Met        | Val        | Leu        |
| Val        | Asn<br>530 | Ala         | Met        | Glu        | Met        | Ala<br>535 | Arg        | Val        | Thr         | Gly         | Va I<br>540 | Pro        | Leu        | Gly        | Tyr        |
| Leu<br>545 | Leu        | Ser         | Arg        | Gly        | GIn<br>550 | Gln        | Val        | Lys        | Val         | Va I<br>555 | Ser         | Gln        | Leu        | Leu        | Arg<br>560 |
| GIn        | Ala        | Met         | Arg        | GIn<br>565 | Gly        | Leu        | Leu        | Met        | Pro<br>570  | Val         | Val         | Lys        | Thr        | Glu<br>575 | Gly        |
| Gly        | Glu        | Asp         | Tyr<br>580 | Thr        | Gly        | Ala        | Thr        | Val<br>585 | lle         | Glu         | Pro         | Leu        | Lys<br>590 | Gly        | Tyr        |
| Tyr        | Asp        | Va I<br>595 | Pro        | He         | Ala        | Thr        | Leu<br>600 | Asp        | Phe         | Ser         | Ser         | Leu<br>605 | Tyr        | Pro        | Ser        |
| He         | Met<br>610 | Met         | Ala        | His        | Asn        | Leu<br>615 | Cys        | Tyr        | Thr         | Thr         | Leu<br>620  | Leu        | Arg        | Pro        | Gly        |
| Ala<br>625 | Ala        | GIn         | Lys        | Leu        | Gly<br>630 | Leu        | Thr        | Glu        | Asp         | GIn<br>635  | Phe         | lle        | Lys        | Thr        | Pro<br>640 |
| Thr        | Gly        | Asp         | Glu        | Phe<br>645 | Val        | Lys        | Ala        | Ser        | Va I<br>650 | Arg         | Lys         | Gly<br>·   | Leu        | Leu<br>655 | Pro        |
| GIn        | lle        | Leu         | Glu<br>660 | Asn        | Leu        | Leu        | Ser        | Ala<br>665 | Arg         | Lys         | Arg         | Ala        | Lys<br>670 | Ala        | Glu        |
| Leu        | Ala        | Lys<br>675  | Glu        | Thr        | Asp        | Pro        | Leu<br>680 | Arg        | Arg         | Gin         | Val         | Leu<br>685 | Asp        | Gly        | Arg        |
| Gln        | Leu        | Ala         | Leu        | Lys        | Val        | Ser        | Ala        | Asn        | Ser         | Val         | Tyr         | Gly        | Phe        | Thr        | Gly        |

|            | 690        |            |             |            |            | 695        |            |            |            |             | 700              |            |             |            |            |
|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------------|------------|-------------|------------|------------|
| Ala<br>705 | GIn        | Val        | Gly         | Arg        | Leu<br>710 | Pro        | Cys        | Leu        | Glu        | l le<br>715 | Ser <sub>.</sub> | Gln        | Ser         | Val        | Thr<br>720 |
| Gly        | Phe        | Gly        | Arg         | GIn<br>725 | Met        | He         | Glu        | Lys        | Thr<br>730 | Lys         | Gin              | Leu        | Val         | Glu<br>735 | Thr        |
| Lys        | Tyr        | Thr        | Va I<br>740 | Glu        | Asn        | Gly        | Tyr        | Ser<br>745 | Thr        | Ser         | Ala              | Lys        | Va I<br>750 | Val        | Tyr        |
| Gly        | Asp        | Thr<br>755 | Asp         | Ser        | Val        | Met        | Cys<br>760 | Arg        | Phe        | Gly         | Val              | Ser<br>765 | Ser         | Val        | Ala        |
| Glu        | Ala<br>770 | Met        | Ala         | Leu        | Gly        | Arg<br>775 | Glu        | Ala        | Ala        | Asp         | Trp<br>780       | Val        | Ser         | Gly        | His        |
| Phe<br>785 | Pro        | Ser        | Pro         | He         | Arg<br>790 | Leu        | Glu        | Phe        | Glu        | Lys<br>795  | Val              | Tyr        | Phe         | Pro        | Tyr<br>800 |
| Leu        | Leu        | He         | Ser         | Lys<br>805 | Lys        | Arg        | Tyr        | Ala        | Gly<br>810 | Leu         | Leu              | Phe        | Ser         | Ser<br>815 | Arg        |
| Pro        | Asp        | Ala        | His<br>820  | Asp        | Arg        | Met        | Asp        | Cys<br>825 | Lys        | Gly         | Leu              | Glu        | Ala<br>830  | Val        | Arg        |
| Arg        | Asp        | Asn<br>835 | Cys         | Pro        | Leu        | Val        | Ala<br>840 | Asn        | Leu        | Val         | Thr              | Ala<br>845 | Ser         | Leu        | Arg        |
| Arg        | Leu<br>850 | Leu        | lle         | Asp        | Arg        | Asp<br>855 | Pro        | Ser        | Gly        | Ala         | Val<br>860       | Ala        | His         | Ala        | Gin        |
| Asp<br>865 | Val        | lle        | Ser         | Asp        | Leu<br>870 | Leu        | Cys        | Asn        | Arg        | l le<br>875 | Asp              | He         | Ser         | Gln        | Leu<br>880 |
| Val        | lle        | Thr        | Lys         | Glu<br>885 | Leu        | Thr        | Arg        | Ala        | Ala<br>890 | Ala         | Asp              | Tyr        | Ala         | Gly<br>895 | Lys        |

| Gin Ala His         | Val Glu<br>900  | Leu Ala         |               | rg Met .<br>05 | Arg Lys         | Arg Asp<br>910  | Pro Gly         |
|---------------------|-----------------|-----------------|---------------|----------------|-----------------|-----------------|-----------------|
| Ser Ala Pro<br>915  | Ser Leu         | Gly Asp         | Arg Va<br>920 | al Pro         | Tyr Val         | lle lle<br>925  | Ser Ala         |
| Ala Lys Gly<br>930  | Val Ala         | Ala Tyr<br>935  | Met L         | ys Ser         | Glu Asp<br>940  | Pro Leu         | Phe Val         |
| Leu Glu His<br>945  | Ser Leu         | Pro IIe         | Asp T         |                | Tyr Tyr<br>955  | Leu Glu         | Gin Gin<br>960  |
| Leu Ala Lys         | Pro Leu<br>965  | Leu Arg         | lle P         | he Glu<br>970  | Pro lle         | Leu Gly         | Glu Gly<br>975  |
| Arg Ala Glu         | Ala Val         | Leu Leu         |               | ily Asp<br>85  | His Thr         | Arg Cys<br>990  | Lys Thr         |
| Val Leu Thr<br>995  |                 |                 | Gly L<br>1000 | eu Leu         |                 | Ala Lys<br>1005 | Arg Arg         |
| Asn Cys Cys<br>1010 | lle Gly         | Cys Arg<br>1015 |               | 'al Leu        | Ser His<br>1020 | Gin Giy         | Ala Val         |
| Cys Lys Phe         |                 | Pro Arg<br>1030 | Glu S         |                | Leu Tyr<br>035  | Gin Lys         | Glu Val<br>1040 |
| Ser His Leu         | Ser Ala<br>1045 |                 | Glu A         | rg Phe<br>1050 | Ser Arg         |                 | Thr Gin<br>1055 |
| Cys Gin Arg         | Cys Gin<br>1060 | Gly Ser         |               | lis Glu<br>165 | Asp Val         | lle Cys<br>1070 | Thr Ser         |
| Arg Asp Cys         |                 |                 | Met A<br>1080 | arg Lys        |                 | Arg Lys<br>1085 | Asp Leu         |

Glu Asp Gln Glu Arg Leu Leu Arg Arg Phe Gly Pro Pro Gly Pro Glu 1090 1095 1100

Ala Trp 1105

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<212> DNA

⟨213⟩ Drosophila melanogaster

#### <400> 63

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|------------|------------|------------|-------------|------------|-------------|------|
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| ggaactcgtg | agaagctgcg | gcagcaggag | aacctgcagg  | acgatcaagt | ggaacgtacg  | 1980 |
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| _             | > 10 |           |      |             |          |             |     |            |      |             |      |     |         |           |      |
|---------------|------|-----------|------|-------------|----------|-------------|-----|------------|------|-------------|------|-----|---------|-----------|------|
|               | > PR |           |      |             |          |             |     |            |      |             |      |     |         |           |      |
| (213          | > Dr | osop      | hila | mel         | anog     | aste        | r   |            |      |             |      |     |         |           |      |
| <4 <b>0</b> 0 | > 64 | Ļ         |      |             |          |             |     |            |      |             |      |     |         |           |      |
| Met           | Asp  | Gly       | Lys  | Arg         | Lys      | Phe         | Asn | Gly        | Thr  | Ser         | Asn  | Gly | His     | Ala       | Lys  |
| 1             |      |           |      | 5           |          |             |     |            | 10   |             |      |     |         | 15        |      |
| Lys           | Pro  | Arg       | Asn  | Pro         | Asp      | Asp         | Asp | Glu        | Glu  | Met         | Gly  | Phe | Glu     | Ala       | Glu  |
| -             |      |           | 20   |             |          |             |     | 25         |      |             |      |     | 30      |           |      |
|               |      |           | D)   | <b>0</b> 1. | <b>.</b> | <b>0</b>    | 01  | <b>.</b>   | Mada | <b>A</b>    | ٥١   | 772 | 1       | 1         | Wa+  |
| Leu           | Ala  | A1a<br>35 | rne  | GIU         | ASN      | ser         | 40  | ASP        | WET  | ASP         | GIN  | 45  | Leu     | Leu       | WEL  |
|               |      |           |      |             |          |             |     |            |      |             |      |     |         |           |      |
| Gly           | Asp  | Gly       | Pro  | Glu         | Asn      | Gln         | Thr | Thr        | Ser  | Glu         | Arg  | Trp | Ser     | Arg       | Pro  |
|               | 50   |           |      |             |          | . <b>55</b> |     |            |      |             | 60   |     |         |           |      |
| Pro           | Pro  | Pro       | Glu  | Leu         | Asp      | Pro         | Ser | Lys        | His  | Asn         | Leu  | Glu | Phe     | Gln       | Gln  |
| 65            |      |           |      |             | 70       |             |     |            |      | 75          |      |     |         |           | 80   |
| _             |      |           | •    |             | <b>.</b> | •           | 01  | 01         | D    | 1           | Dan  | 01  | 1# - J. | Dun       | Cl., |
| Leu           | Asp  | Val       | Glu  | Asn<br>85   | lyr      | Leu         | GIY | GIN        | 90   | Leu         | Pro  | uly | WET     | Pro<br>95 | шу   |
|               |      |           |      |             |          |             |     |            |      | ,           |      |     |         |           |      |
| Ala           | Gln  | He        | Gly  | Pro         | Val      | Pro         | Val | Val        | Arg  | Met         | Phe  | Gly |         | Thr       | Met  |
|               |      |           | 100  |             |          |             | ٠   | 105        |      |             |      |     | 110     |           |      |
| Glu           | Gly  | Asn       | Ser  | Val         | Cys      | Cys         | His | Val        | His  | Gly         | Phe  | Cys | Pro     | Tyr       | Phe  |
|               |      | 115       |      |             |          |             | 120 |            |      |             |      | 125 |         |           |      |
| _             |      | 01        | 41.  |             | 0        | <b>0</b> 1  | Di  | <b>C</b> I | CI   | u: <b>-</b> | u: - | ٥   | Cl.,    | 1.40      | Lau  |
| ıyr           | 11e  |           | Ala  | Pro         | ser      | 135         |     | uiu        | diu  | шѕ          | 140  | Uys | diu     | Lys       | Leu  |
|               |      |           |      |             |          |             |     |            |      |             |      |     |         |           |      |
| Gln           | Lys  | Ala       | Leu  | Asp         | Gln      | Lys         | Val | lle        | Ala  | Asp         | He   | Arg | Asn     | Asn       |      |
| 145           |      |           |      |             | 150      |             |     |            |      | 155         |      |     |         |           | 160  |
| Asp           | Asn  | Val       | Gln  | Glu         | Ala      | Val         | Leu | Met        | Val  | Glu         | Leu  | Val | Glu     | Lys       | Leu  |
|               |      | •         | •    | 165         |          |             |     |            | 170  |             |      |     |         | 175       |      |

| Asn        | lle        | His        | Gly<br>180 | Tyr         | Asn        | Gly        | Asp         | Lys<br>185 | Lys        | Gln         | Arg        |            | 11e<br>190 | Lys         | He          |
|------------|------------|------------|------------|-------------|------------|------------|-------------|------------|------------|-------------|------------|------------|------------|-------------|-------------|
| Ser        | Vai        | Thr<br>195 | Leu        | Pró         | Arg        | Phe        | Va I<br>200 | Ala        | Ala        | Ala         | Ser        | Arg<br>205 | Leu        | Leu         | Lys         |
| Lys        | Glu<br>210 | Val        | lle        | Met         | Ser        | Glu<br>215 | He          | Asp        | Phe        | Gln         | Asp<br>220 | Cys        | Arg        | Ala         | Phe         |
| Glu<br>225 | Asn        | Asn        | He         | Asp         | Phe<br>230 | Asp        | lle         | Arg        | Phe        | Met<br>235  | Val        | Asp        | Thr        | Asp         | Va i<br>240 |
| Val        | Gly        | Cys        | Asn        | Trp<br>245  | lle        | Glu        | Leu         | Pro        | Met<br>250 | Gly         | His        | Trp        | Arg        | l le<br>255 | Arg         |
| Asn        | Ser        | His        | Ser<br>260 | Lys         | Pro        | Leu        | Pro         | Glu<br>265 | Ser        | Arg         | Cys        | Gin        | lle<br>270 | Glu         | Val         |
| Asp        | Val        | Ala<br>275 |            | Asp         | Arg        | Phe        | 11e<br>280  | Ser        | His        | Glu         | Pro        | Glu<br>285 | Gly        | Glu         | Trp         |
| Ser        | Lys<br>290 | Val        | Ala        | Pro         | Phe        | Arg<br>295 | He          | Leu        | Ser        | Phe         | Asp<br>300 | lle        | Glu        | Cys         | Ala         |
| Gly<br>305 | Arg        | Lys        | Gly        | He          | Phe<br>310 | Pro        | Glu         | Ala        | Lys        | i le<br>315 | Asp        | Pro        | Vaİ        | lle         | GIn<br>320  |
| lle        | Ala        | Asn        | Met        | Va I<br>325 |            | Arg        | Gin         | Gly        | Glu<br>330 | Arg         | Glu        | Pro        | Phe        | 11e<br>335  | Arg         |
| Asn        | Val        | Phe        | Thr<br>340 |             | Asn        | Glu        | Cys         | Ala<br>345 |            | lle         | He         | Gly        | Ser<br>350 | GIn         | Val         |
| Leu        | Cys        | His<br>355 |            | Lys         | Glu        | Thr        | GIn<br>360  | Met        | Leu        | Asp         | Lys        | Trp<br>365 |            | Aia         | Phe         |

| Val        | Arg<br>370 | Glu        | Val        | Asp        | Pro        | Asp<br>375 | lle        | Ļeu        | Thr        | Gly        | Tyr<br>380 | Asn         | lle        | Asn         | Asn        |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|-------------|------------|
| Phe<br>385 | Asp        | Phe        | Pro        | Tyr        | Leu<br>390 | Leu        | Asn        | Arg        | Ala        | Ala<br>395 | His        | Leu         | Lys        | Val         | Arg<br>400 |
| Asn        | Phe        | Glu        | Tyr        | Leu<br>405 | Gly        | Arg        | He         | Lys        | Asn<br>410 | He         | Arg        | Ser         | Val        | l le<br>415 | Lys        |
| Glu        | Gln        | Met        | Leu<br>420 | Gln        | Ser        | Lys        | Gln        | Met<br>425 | Gly        | Arg        | Arg        | Glu         | Asn<br>430 | GIn         | Tyr        |
| Val        | Asn        | Phe<br>435 | Glu        | Gly        | Arg        | Val        | Pro<br>440 | Phe        | Asp        | Leu        | Leu        | Phe<br>445  | Val        | Leu         | Leu        |
| Arg        | Asp<br>450 | Tyr        | Lys        | Leu        | Arg        | Ser<br>455 | Tyr        | Thr        | Leu        | Asn        | Ala<br>460 | Val         | Ser        | Tyr         | His        |
| Phe<br>465 | Leu<br>·   | Gin        | Glu        | Gln        | Lys<br>470 | Glu        | Asp        | Val        | His        | His<br>475 | Ser        | He          | lle        | Thr         | Asp<br>480 |
| Leu        | Gin        | Asn        | Gly        | Asp<br>485 | Glu        | Gin        | Thr        | Arg        | Arg<br>490 | Arg        | Ser        | Ala         | Met        | Tyr<br>495  | Cys        |
| Leu        | Lys        | Asp        | Ala<br>500 | Tyr        | Leu        | Pro        | Leu        | Arg<br>505 | Leu        | Leu        | Glu        | Lys         | Leu<br>510 | Met         | Ala        |
| lle        | Val        | Asn<br>515 | Tyr        | Met        | Glu        | Met        | Ala<br>520 | Arg        | Val        | Thr        | Gly        | Va I<br>525 | Pro        | Leu         | Glu        |
| Ser        | Leu<br>530 |            | Thr        | Arg        | Gly        | GIn<br>535 | Gln        | lle        | Lys        | Val        | Leu<br>540 | Ser         | Gin        | Leu         | Leu        |
| Arg<br>545 | Lys        | Ala        | Lys        | Thr        | Lys<br>550 | Gly        | Phe        | lle        | Met        | Pro<br>555 | Ser        | Tyr         | Thr        | Ser         | GIn<br>560 |
| Gly        | Ser        | Asp        | Glu        | Gln        | Tyr        | Glu        | Gly        | Ala        | Thr        | Val        | He         | Glu         | Pro        | Lys         | Arg        |

|             |            |            |             | 565        |            |            |            |             | 570        |            |            |            |             | 575        |             |
|-------------|------------|------------|-------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|-------------|------------|-------------|
| Gly         | Tyr        | Tyr        | Ala<br>580  | Asp        | Pro        | He         | Ser        | Thr<br>585  | Leu        | Asp        | Phe        | Ala        | Ser<br>590  | Leu        | Tyr         |
| Pro         | Ser        | lle<br>595 | Met         | Met        | Ala        | His        | Asn<br>600 | Leu         | Cys        | Tyr        | Thr        | Thr<br>605 | Leu         | Val        | Leu         |
| Gly         | Gly<br>610 | Thr        | Arg         | Glu        | Lys        | Leu<br>615 | Arg        | Gln         | GIn        | Glu        | Asn<br>620 | Leu        | Gln         | Asp        | Asp         |
| G1n<br>625  | Val        | Glu        | Arg         | Thr        | Pro<br>630 | Ala        | Asn        | Asn         | Tyr        | Phe<br>635 | Val        | Lys        | Ser         | Glu        | Va I<br>640 |
| Arg         | Arg        | Gly        | Leu         | Leu<br>645 | Pro        | Glu        | lle        | Leu         | Glu<br>650 | Ser        | Leu        | Leu        | Ala         | Ala<br>655 | Arg         |
| Lys         | Arg        | Ala        | Lys<br>660  | Asn        | Asp        | Leu        | Lys        | Va I<br>665 | Glu        | Thr        | Asp        | Pro        | Phe<br>670  | Lys        | Arg         |
| Lys         | Val        | Leu<br>675 | Asp         | Gly        | Arg        | Gln        | Leu<br>680 | Ala         | Leu        | Lys        | He         | Ser<br>685 | Ala         | Asn        | Ser         |
| Val         | Tyr<br>690 | Gly        | Phe         | Thr        | Gly        | Ala<br>695 | Gin        | Val         | Gly        | Lys        | Leu<br>700 | Pro        | Cys         | Leu        | Glu         |
| l le<br>705 | Ser        | Gly        | Ser         | Val        | Thr<br>710 | Ala        | Tyr        | Gly         | Arg        | Thr<br>715 | Met        | lle        | Glu         | Met        | Thr<br>720  |
| Lys         | Asn        | Glu        | Val         | Glu<br>725 |            | His        | Tyr        | Thr         | GIn<br>730 | Ala        | Asn        | Gly        | Tyr         | Glu<br>735 | Asr         |
| Asn         | Ala        | Val        | Va I<br>740 |            | Tyr        | Gly        | Asp        | Thr<br>745  | Asp        | Ser        | Val        | Met        | Va I<br>750 | Asn        | Phe         |
| Gly         | Val        | Lys<br>755 | Thr         | Leu        | Glu        | Arg        | Ser<br>760 | Met         | Glu        | Leu        | Gly        | Arg<br>765 | Glu         | Ala        | Ala         |

| Glu        | Leu<br>770 | Val        | Ser         | Ser        | Lys         | Phe<br>775 | Val        | His        | Pro        | lle        | Lys<br>780 <sub>.</sub> |            | Glu        | Phe        | Glu        |
|------------|------------|------------|-------------|------------|-------------|------------|------------|------------|------------|------------|-------------------------|------------|------------|------------|------------|
| Lys<br>785 | Vai        | Tyr        | Tyr         | Pro        | Tyr<br>790  | Leu        | Leu        | lle        | Asn        | Lys<br>795 | Lys                     | Arg        | Tyr        | Ala        | Gly<br>800 |
| Leu        | Tyr        | Phe        | Thr         | Arg<br>805 | Pro         | Asp        | Thr        | Tyr        | Asp<br>810 | Lys        | Met                     | Asp        | Cys        | Lys<br>815 | Gly        |
| He         | Glu        | Thr        | Va I<br>820 | Arg        | Arg         | Asp        | Asn        | Ser<br>825 | Pro        | Leu        | Val                     | Ala        | Asn<br>830 | Leu        | Met        |
| Asn        | Ser        | Cys<br>835 | Leu         | Gin        | Lys         | Leu        | Leu<br>840 | He         | Glu        | Arg        | Asp                     | Pro<br>845 | Asp        | Gly        | Ala        |
| Val        | Ala<br>850 | Tyr        | Val         | Lys        | GIn         | Val<br>855 | lle        | Ala        | Asp        | Leu        | Leu<br>860              | Cys        | Asn        | Arg        | lle        |
| Asp<br>865 | lle        | Ser        | His         | Leu        | Va I<br>870 | He         | Thr        | Lys        | Glu        | Leu<br>875 | Ala                     | Lys        | Thr        | Asp        | Tyr<br>880 |
| Ala        | Ala        | Lys        | Gln         | Ala<br>885 | His         | Val        | Glu        | Leu        | A1a<br>890 | Ala        | Lys                     | Met        | Lys        | Lys<br>895 | Arg        |
| Asp        | Pro        | Gly        | Thr<br>900  | Ala        | Pro         | Lys        | Leu        | Gly<br>905 | Asp        | Arg        | Val                     | Pro        | Tyr<br>910 | Val        | He         |
| Cys        | Ala        | Ala<br>915 |             | Lys        | Asn         | Thr        | Pro<br>920 | Ala        | Tyr        | Gln        | Lys<br>·                | Ala<br>925 | Glu        | Asp        | Pro        |
| Leu        | Tyr<br>930 |            | Leu         | Glu        | Asn         | Ser<br>935 | Val        | Pro        | lle        | Asp        | Ala<br>940              | Thr        | Tyr        | Tyr        | Leu        |
| Glu<br>945 |            | Gln        | Leu         | Ser        | Lys<br>950  |            | Leu        | Leu        | Arg        | lle<br>955 | Phe                     | Glu        | Pro        | He         | Leu<br>960 |

Gly Asp Asn Ala Glu Ser IIe Leu Leu Lys Gly Glu His Thr Arg Thr 965 970 975

Arg Thr Val Val Thr Ser Lys Val Gly Gly Leu Ala Gly Phe Met Thr 980 985 990

Lys Lys Thr Ser Cys Leu Gly Cys Lys Ser Leu Met Pro Lys Gly Tyr 995 1000 1005

Giu Gin Ala Cys Leu Cys Pro His Cys Giu Pro Arg Met Ser Giu Leu 1010 1015 1020

Tyr Gin Lys Giu Vai Gly Ala Lys Arg Giu Leu Giu Giu Thr Phe Ser 1025 1030 1035 1040

Arg Leu Trp Thr Glu Cys Gln Arg Cys Gln Glu Ser Leu His Glu Glu

1045 1050 1055

Val lie Cys Ser Asn Arg Asp Cys Pro lie Phe Tyr Met Arg Gln Lys 1060 1065 1070

Val Arg Met Asp Leu Asp Asn Gln Glu Lys Arg Val Leu Arg Phe Gly 1075 1080 1085

Leu Ala Glu Trp 1090

<210> 65

<211> 9064

<212> DNA

<213> Drosophila melanogaster

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| tatggaactg  | aaacgactgc | aatgttttta   | tattccggca | acgcatttcg  | cataaattac | 240  |
|-------------|------------|--------------|------------|-------------|------------|------|
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| cagaaaaaagg | gcaataagat | tcgggataac   | aaaacttgat | ggcgttcccg  | attttcccgg | 360  |
| acaagggagc  | gtatatgtat | gtacacacaa   | aaaaaaaact | taaccagcct  | tgcataacga | 420  |
| aacacgtgca  | ataaaaatat | gactgattgt   | caacctctgc | tgcaacttaa  | ttgctgccgc | 480  |
| agaggtaaaa  | ctgaaaaaca | taaaaggggg   | gcgacaagtg | cagcaagcga  | aaaaataaag | 540  |
| aactcaaaga  | gcgcatgcgc | gccgctctcc   | cactctctct | ccctctctct  | ctgtcgctcc | 600  |
| actgcgctgg  | aatcttacaa | ctcgtgtagg   | tgagccggat | ttttatgatg  | atgcgcctgt | 660  |
| gtgcgtgact  | gcatgatgcc | attgcagcgg   | agaactagta | gaaaaaaagtt | cacatttcag | 720  |
| cagttggaaa  | acacatggcc | aacaggccaa   | ctcaagtggc | cagcagctgt  | ccttatattg | 780  |
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|             |            | ttgtattgtg   |            | •           |            |      |
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| Į | ggtgttttcc | gggcggacat   | accatgccgt | tttcgtctag   | atcctgctat   | ggtcaagcaa | 3000 |
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|   |            |              |            |              | tggtgttacc   |            |      |
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|   | _          |              |            |              | a gaggatcaaa |            |      |
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9064

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| \$\frac{210}{66}  |                 |       | 5 C L 1    | 56 -6  | LLAB         | Lg a       | -e -e,      | POP | t Bu | -666 | LELQ        | rgo | saca | JLE  | COBL | асвищ |
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| <213> Drosophila melanogaster         <400> 66         Met Ser Asp Ser Gly Lys Gly Lys Val Leu Gln Asn Thr Gly Lys Phe 1  | <b>&lt;21</b> 1 | > 22  | 220        |        |              |            |             |     |      |      |             |     |      |      |      |       |
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| Met         Ser         Asp         Ser         Gly         Lys         Gly         Lys         Val         Leu         Gln         Asn         Thr         Gly         Lys         Phe         Phe         Asn         Gly         Lys         Phe         Phe         Asn         Glu         Ala         Gly         Asp         Phe         Phe         Asn         Glu         Ala         Gly         Phe         Phe         Asn         Glu         Ala         Gly         Phe         Phe         Asn         Glu         Ala         Ala         Leu         Ala         Ala         Leu         Ala         Ala <td>&lt;213</td> <td>3&gt; Di</td> <td>roso</td> <td>ohi la</td> <td>a me</td> <td>anog</td> <td>gaste</td> <td>er</td> <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> | <213            | 3> Di | roso       | ohi la | a me         | anog       | gaste       | er  |      |      |             |     |      |      |      |       |
| Met         Ser         Asp         Ser         Gly         Lys         Gly         Lys         Val         Leu         Gln         Asn         Thr         Gly         Lys         Phe         Phe         Asn         Gly         Lys         Phe         Phe         Asn         Glu         Ala         Gly         Asp         Phe         Phe         Asn         Glu         Ala         Gly         Phe         Phe         Asn         Glu         Ala         Gly         Phe         Phe         Asn         Glu         Ala         Ala         Leu         Ala         Ala         Leu         Ala         Ala <td></td>   |                 |       |            |        |              |            |             |     |      |      |             |     |      |      |      |       |
| 1 5 10 15  Val Ser Glu Asn Arg Thr Glu Gly Asp Asp Phe Phe Asn Glu Ala Gly 20 25 30  Tyr Arg Gln Ser Arg Glu Asn Asp Lys I le Asp Ser Lys Tyr Gly Phe 35 40 45  Asp Arg Val Lys Asp Ser Gln Glu Arg Thr Gly Tyr Leu I le Asn Met 50 55 60  His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu I le Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe I le Gln Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gln Pro Tyr Leu Leu I le Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln I le Ser Gly 115 120 125  Leu Glu His I le Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser  | <400            | )> 66 | 3          |        |              |            |             |     |      |      |             |     |      |      |      |       |
| Val Ser Glu Asn Arg Thr Glu Gly Asp Asp Phe Phe Asn Glu Ala Gly 20       25       30         Tyr Arg Gln Ser Arg Glu Asn Asp Lys Ile Asp Ser Lys Tyr Gly Phe 35       40       45         Asp Arg Val Lys Asp Ser Gln Glu Arg Thr Gly Tyr Leu Ile Asn Met 50       55       60         His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu Ile Ala Ala Leu 65       70       75       80         Asp Leu Phe Phe Ile Gln Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85       90       95         Ala Tyr Gln Pro Tyr Leu Leu Ile Arg Pro Glu Asp Asn Met His Leu 100       105       110         Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln Ile Ser Gly 115       120       125         Leu Glu His Ile Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser       125   | Met             | Ser   | Asp        | Ser    | Gly          | Lys        | Gly         | Lys | Val  | Leu  | Gln         | Asn | Thr  | Gly  | Lys  | Phe   |
| 20 25 30  Tyr Arg Gin Ser Arg Giu Asn Asp Lys IIe Asp Ser Lys Tyr Giy Phe 35 40 45  Asp Arg Val Lys Asp Ser Gin Giu Arg Thr Giy Tyr Leu IIe Asn Met 50 55 60  His Ser Asn Giu Val Leu Asp Giu Asp Arg Arg Leu IIe Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe IIe Gin Met Asp Giy Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu IIe Arg Pro Giu Asp Asn Met His Leu 100 105 110  Giu Val Ala Arg Phe Leu Giy Arg Lys Tyr Ser Giy Gin IIe Ser Giy 115 120 125  Leu Giu His IIe Thr Lys Giu Asp Leu Asp Leu Pro Asn His Leu Ser  | 1               |       |            |        | 5            |            |             |     |      | 10   |             |     |      |      | 15   |       |
| 20 25 30  Tyr Arg Gin Ser Arg Giu Asn Asp Lys IIe Asp Ser Lys Tyr Giy Phe 35 40 45  Asp Arg Val Lys Asp Ser Gin Giu Arg Thr Giy Tyr Leu IIe Asn Met 50 55 60  His Ser Asn Giu Val Leu Asp Giu Asp Arg Arg Leu IIe Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe IIe Gin Met Asp Giy Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu IIe Arg Pro Giu Asp Asn Met His Leu 100 105 110  Giu Val Ala Arg Phe Leu Giy Arg Lys Tyr Ser Giy Gin IIe Ser Giy 115 120 125  Leu Giu His IIe Thr Lys Giu Asp Leu Asp Leu Pro Asn His Leu Ser  |                 | _     |            | _      | _            | _          |             |     | _    |      |             |     | _    |      |      |       |
| Tyr Arg Gin Ser Arg Giu Asn Asp Lys I le Asp Ser Lys Tyr Giy Phe 35 40 45  Asp Arg Val Lys Asp Ser Gin Giu Arg Thr Giy Tyr Leu I le Asn Met 50 55 60  His Ser Asn Giu Val Leu Asp Giu Asp Arg Arg Leu I le Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe I le Gin Met Asp Giy Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu I le Arg Pro Giu Asp Asn Met His Leu 100 105 110  Giu Val Ala Arg Phe Leu Giy Arg Lys Tyr Ser Giy Gin I le Ser Giy 115 120 125  Leu Giu His I le Thr Lys Giu Asp Leu Asp Leu Pro Asn His Leu Ser   | Val             | Ser   | Glu        |        | Arg          | Thr        | Glu         | Gly |      | Asp  | Phe         | Phe | Asn  |      | Ala  | Gly   |
| Asp Arg Val Lys Asp Ser Gin Giu Arg Thr Gly Tyr Leu lie Asn Met 50 55 60   His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu lie Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe lie Gin Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu lie Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gin lie Ser Gly 115 120 125  Leu Glu His lie Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   |                 |       |            | 20     |              |            |             |     | 25   |      |             |     |      | 30   | •    |       |
| Asp Arg Val Lys Asp Ser Gin Giu Arg Thr Gly Tyr Leu lie Asn Met 50 55 60   His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu lie Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe lie Gin Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu lie Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gin lie Ser Gly 115 120 125  Leu Glu His lie Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   | Tvr             | Δrσ   | Gln        | Ser    | Δrσ          | Glu        | <b>∆</b> en | Δen | lve  | ماا  | <b>A</b> en | Ser | l ve | Tvr  | Gly  | Phe   |
| Asp Arg Val Lys Asp Ser Gln Glu Arg Thr Gly Tyr Leu IIe Asn Met 50 55 60  His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu IIe Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe IIe Gln Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gln Pro Tyr Leu Leu IIe Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln IIe Ser Gly 115 120 125  Leu Glu His IIe Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser  |                 | / u B |            | 00.    | / <b>u</b> B | <b>u.u</b> |             |     | _,0  |      | nop         | 00. |      | . ,. | u.,  |       |
| His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu IIe Ala Ala Leu 65 70 75 80  Asp Leu Phe Phe IIe Gln Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gln Pro Tyr Leu Leu IIe Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln IIe Ser Gly 115 120 125  Leu Glu His IIe Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser  |                 |       |            |        |              |            |             |     |      |      |             |     | -    |      |      |       |
| His Ser Asn Glu Val Leu Asp Glu Asp Arg Arg Leu IIe Ala Ala Leu 70 75 80  Asp Leu Phe Phe IIe Gln Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gln Pro Tyr Leu Leu IIe Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln IIe Ser Gly 115 120 125  Leu Glu His IIe Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   | Asp             | Arg   | Val        | Lys    | Asp          | Ser        | Gln         | Glu | Arg  | Thr  | Gly         | Tyr | Leu  | lle  | Asn  | Met   |
| Asp Leu Phe Phe lle Gin Met Asp Giy Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu lle Arg Pro Giu Asp Asn Met His Leu 100 105 110  Giu Val Ala Arg Phe Leu Giy Arg Lys Tyr Ser Giy Gin Ile Ser Giy 115 120 125  Leu Giu His Ile Thr Lys Giu Asp Leu Asp Leu Pro Asn His Leu Ser   |                 | 50    |            |        |              |            | 55          |     |      |      |             | 60  |      |      |      |       |
| Asp Leu Phe Phe lle Gin Met Asp Giy Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu lle Arg Pro Giu Asp Asn Met His Leu 100 105 110  Giu Val Ala Arg Phe Leu Giy Arg Lys Tyr Ser Giy Gin Ile Ser Giy 115 120 125  Leu Giu His Ile Thr Lys Giu Asp Leu Asp Leu Pro Asn His Leu Ser   |                 |       |            |        |              |            |             |     |      |      |             |     |      |      |      |       |
| Asp Leu Phe Phe lie Gin Met Asp Gly Ser Arg Phe Lys Cys Thr Val 85 90 95  Ala Tyr Gin Pro Tyr Leu Leu lie Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gin lie Ser Gly 115 120 125  Leu Glu His lie Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   | His             | Ser   | Asn        | Glu    | Val          | Leu        | Asp         | Glu | Asp  | Arg  | Arg         | Leu | He   | Ala  | Ala  | Leu   |
| Ala Tyr Gln Pro Tyr Leu Leu IIe Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln IIe Ser Gly 115 120 125  Leu Glu His IIe Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   | 65              |       |            |        |              | 70         |             | •   |      |      | 75          |     |      |      |      | 80    |
| Ala Tyr Gln Pro Tyr Leu Leu IIe Arg Pro Glu Asp Asn Met His Leu 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln IIe Ser Gly 115 120 125  Leu Glu His IIe Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   |                 |       | <b>D</b> 1 | -      |              | 01         |             |     |      | _    |             | ъ.  |      | •    | ~    |       |
| Ala Tyr Gin Pro Tyr Leu Leu I le Arg Pro Giu Asp Asn Met His Leu 100 105 110  Giu Val Ala Arg Phe Leu Giy Arg Lys Tyr Ser Giy Gin I le Ser Giy 115 120 125  Leu Giu His I le Thr Lys Giu Asp Leu Asp Leu Pro Asn His Leu Ser  | Asp             | Leu   | Phe        | rne    |              | GIN        | Met         | Asp | GIY  |      | Arg         | rne | Lys  | Cys  |      | Val   |
| 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln lle Ser Gly 115 120 125  Leu Glu His Ile Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   |                 |       |            |        | 65           |            |             |     |      | 90   |             |     |      |      | 90   |       |
| 100 105 110  Glu Val Ala Arg Phe Leu Gly Arg Lys Tyr Ser Gly Gln lle Ser Gly 115 120 125  Leu Glu His Ile Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   | Ala             | Tvr   | Gln        | Pro    | Tvr          | Leu        | Leu         | He  | Arg  | Pro  | Glu         | Asp | Asn  | Met  | His  | Leu   |
| 115 120 125<br>Leu Glu His Ile Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser  |                 |       |            |        |              |            |             |     |      |      |             |     |      |      |      |       |
| 115 120 125<br>Leu Glu His Ile Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser  |                 |       |            |        |              |            |             |     |      |      |             | -   |      |      |      |       |
| Leu Glu His Ile Thr Lys Glu Asp Leu Asp Leu Pro Asn His Leu Ser   | Glu             | Val   | Ala        | Arg    | Phe          | Leu        | Gly         | Arg | Lys  | Tyr  | Ser         | Gly | Gln  | He   | Ser  | Gly   |
|   |                 |       | 115        |        |              |            |             | 120 |      |      |             |     | 125  |      |      |       |
|   |                 |       |            |        |              |            |             |     |      |      |             |     |      |      |      |       |
| 400   | Leu             |       | His        | He     | Thr          | Lys        |             | Asp | Leu  | Asp  | Leu         |     | Asn  | His  | Leu  | Ser   |
| 130 135 140   |                 | 130   |            |        |              |            | 135         |     |      |      |             | 140 |      |      |      |       |

| Gly Le        | eu Gin               | GIn        | GIn        | Tyr<br>150  | lle        | Lys        | Leu        | Ser         | Phe<br>155 | Leu        | Asn        | Gln        | Thr         | Aia<br>160 |
|---------------|----------------------|------------|------------|-------------|------------|------------|------------|-------------|------------|------------|------------|------------|-------------|------------|
| Met Th        | nr Lys               | Val        | Arg<br>165 | Arg         | Glu        | Leu        | Met        | Ser<br>170  | Ala        | Val        | Lys        | Arg        | Asn<br>175  | Gin        |
| Glu Ar        | g Gln                | Lys<br>180 | Ser        | Asn         | Thr        | Tyr        | Tyr<br>185 | Met         | Gln        | Met        | Leu        | Ala<br>190 | Thr         | Ser        |
| Leu A         | la Gin<br>195        |            | Ser        | Ala         | Gly        | Ser<br>200 | Glu        | Asp         | Ala        | Thr        | Leu<br>205 | Gly        | Lys         | Arg        |
|               | In <b>A</b> sp<br>10 | Tyr        | Met        | Asp         | Cys<br>215 |            | Val        | Asp         | lle        | Arg<br>220 | Glu        | His        | Asp         | Val        |
| Pro Ty<br>225 | yr His               | Val        | Arg        | Va I<br>230 | Ser        | lle        | Asp        | Leu         | Arg<br>235 |            | Phe        | Cys        | Gly         | GIn<br>240 |
| Trp T         | yr Asn               | lle        | Arg<br>245 | Cys         | Arg        | Ser        | Gly        | Va I<br>250 | Glu        | Leu        | Pro        | Thr        | l le<br>255 | Thr        |
| Cys A         | rg Pro               | Asp<br>260 | He         | Leu         | Asp        | Arg        | Pro<br>265 | Glu         | Pro        | Val        | Val        | Leu<br>270 | Ala         | Phe        |
| Asp 1         | le Glu<br>275        |            | Thr        | Lys         | Leu        | Pro<br>280 | Leu        | Lys         | Phe        | Pro        | Asp<br>285 | Ala        | Gln         | Thr        |
|               | in Val<br>90         | Met        | Met        | He          | Ser<br>295 |            | Met        | lle         | Asp        | Gly<br>300 | Gln        | Gly        | Tyr         | Leu        |
| lle Ti<br>305 | hr Asn               | Arg        | Glu        | 11e<br>310  | lle        | Ser        | Ser        | Asn         | Vai<br>315 | Asp        | Asp        | Phe        | Glu         | Tyr<br>320 |
| Thr P         | ro Lys               | : Pro      | Glu<br>325 |             | Glu        | Gly        | Asn        | Phe<br>330  | He         | Val        | Phe        | Asn        | Glu<br>335  | Glu        |

| Asn         | Glu         | Met        | Gln<br>340 | Leu        | Leu        | Gln        | Arg        | Phe<br>345  | Phe        | Asp        | His        | lle        | Met<br>350 | Glu        | Vai        |
|-------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|
| Arg         | Pro         | His<br>355 | He         | He         | Va I       | Thr        | Tyr<br>360 | Asn         | Gly        | Asp        | Phe        | Phe<br>365 | Asp        | Trp        | Pro        |
| Phe         | Va I<br>370 | Glu        | Thr        | Arg        | Ala        | Ala<br>375 | Val        | Tyr         | Asp        | Leu        | Asp<br>380 | Met        | Lys        | Gln        | Glu        |
| l le<br>385 | Gly         | Phe        | Ser        | Lys        | Leu<br>390 | Arg        | Asp        | Gly         | Asn        | Tyr<br>395 | Leu        | Ser        | Arg        | Pro        | Ala<br>400 |
| lle         | His         | Met        | Asp        | Cys<br>405 | Leu        | Cys        | Trp        | Val         | Lys<br>410 | Arg        | Asp        | Ser        | Tyr        | Leu<br>415 | Pro        |
| Val         | Gly         | Ser        | GIn<br>420 | Gly        | Leu        | Lys        | Ala        | Va I<br>425 | Ala        | Lys        | Ala        | Lys        | Leu<br>430 | Arg        | Tyr        |
| Asp         | Pro         | Val<br>435 | Glu        | Leu        | Asp        | Pro        | Glu<br>440 | Asp         | Met        | Cys        | Arg        | Met<br>445 | Ala        | Val        | Glu        |
| GIn         | Pro<br>450  | Gln        | Val        | Leu        | Ala        | Asn<br>455 | Tyr        | Ser         | Val        | Ser        | Asp<br>460 | Ala        | Val        | Ala        | Thr        |
| Tyr<br>465  | Tyr         | Leu        | Tyr        | Met        | Lys<br>470 | Tyr        | Val        | His         | Pro        | Phe<br>475 | lle        | Phe        | Ala        | Leu        | Asn<br>480 |
| Thr         | lle         | He         | Pro        | Met<br>485 | Glu        | Pro        | Asp        | Glu         | ile<br>490 | Leu        | Arg        | Lys        | Gly        | Ser<br>495 | Gly        |
| Thr         | Leu         | Cys        | Glu<br>500 | Thr        | Leu        | Leu        | Met        | Val<br>505  | Glu        | Ala        | Tyr        | His        | Ala<br>510 | GIn        | lle        |
| Val         | Tyr         | Pro<br>515 | Asn        | Lys        | His        | Gin        | Ser<br>520 | Glu         | Leu        | Asn        | Lys        | Leu<br>525 | Ser        | Asn        | Glu        |
| Gly         | His         | Val        | Leu        | Asp        | Ser        | Glu        | Thr        | Tyr         | Val        | Gly        | Gly        | His        | Val        | Glu        | Ala        |

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|             | 530        |            |             |            |            | 535        |            |            |            |            | 540              |            |            |            |            |
|-------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|
| Leu<br>545  | Glu        | Ser        | Gly         | Val        | Phe<br>550 | Arg        | Ala        | Asp        | lle        | Pro<br>555 | Cys <sub>.</sub> | Arg        | Phe        | Arg        | Leu<br>560 |
| Asp         | Pro        | Ala        | Met         | Val<br>565 | Lys        | Gin        | Leu        | Gin        | Glu<br>570 | Gln        | Val              | Asp        | Ala        | Val<br>575 | Leu        |
| Arg         | His        | Ala        | l le<br>580 | Glu        | Val        | Glu        | Glu        | Gly<br>585 | He         | Pro        | Leu              | Glu        | Lys<br>590 | Val        | Leu        |
| Asn         | Leu        | Asp<br>595 | Glu         | Val        | Arg        | Gin        | Glu<br>600 | lle        | Val        | Gin        | Gly              | Leu<br>605 | GIn        | Gly        | Leu        |
| His         | Asp<br>610 | He         | Pro         | Asn        | Arg        | Leu<br>615 | Glu        | Gln        | Pro        | Vai        | lle<br>620       | Tyr        | His        | Leu        | Asp        |
| Va I<br>625 | Gly        | Ala        | Met         | Tyr        | Pro<br>630 | Asn        | He         | He         | Leu        | Thr<br>635 | Asn              | Arg        | Leu        | GIn        | Pro<br>640 |
| Ser         | Ala        | Met        | Val         | Ser<br>645 | Asp        | Leu        | Asp        | Cys        | Ala<br>650 | Ala        | Cys              | Asp        | Phe        | Asn<br>655 | Lys        |
| Pro         | Gly        | Val        | Arg<br>660  | Cys        | Lys        | Arg        | Ser        | Met<br>665 | Asp        | Trp        | Leu              | Trp        | Arg<br>670 | Gly        | Glu        |
| Met         | Leu        | Pro<br>675 | Ala         | Ser        | Arg        | Asn        | Glu<br>680 | Phe        | Gin        | Arg        | lle              | GIn<br>685 | Gin        | Gln        | Leu        |
| Glu         | Thr<br>690 | Glu        | Lys         | Phe        | Pro        | Pro<br>695 | Leu        | Phe        | Pro        | Gly        | Gly<br>700       | Pro        | Gln        | Arg        | Ala        |
| Phe<br>705  | His        | Glu        | Leu         | Ser        | Lys<br>710 | Glu        | Asp        | Gin        | Ala        | Ala<br>715 | Tyr              | Glu        | Lys        | Lys        | Arg<br>720 |
| Leu         | Thr        | Asp        | Tyr         | Cys        | Arg        | Lys        | Ala        | Tyr        | Lys        | Lys        | Thr              | Lys        | Leu        | Thr        | Lys        |

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|            |            |            |            | 725        |            |            |            |            | 730        |            |                  |            |            | 735        |             |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------------|------------|------------|------------|-------------|
| Leu        | Glu        | Thr        | Arg<br>740 | Thr        | Ser        | Thr        | lle        | Cys<br>745 | Gln        | Lys        | Glu <sub>.</sub> | Asn        | Ser<br>750 | Phe        | Tyr         |
| Val        | Asp        | Thr<br>755 | Val        | Arg        | Ala        | Phe        | Arg<br>760 | Asp        | Arg        | Arg        | Tyr              | Glu<br>765 | Tyr        | Lys        | Gly         |
| Leu        | Thr<br>770 |            | Val        | Ala        | Lys        | Ala<br>775 | Ser        | Val        | Asn        | Ala        | Ala<br>780       | Val        | Ala        | Ser        | Gly         |
| Asp<br>785 | Ala        | Ala        | Glu        | lle        | Lys<br>790 | Ala        | Ala        | Lys        | Gly        | Arg<br>795 | Glu              | Val        | Leu        | Tyr        | Asp<br>800  |
| Ser        | Leu        | Gln        | Leu        | Ala<br>805 | His        | Lys        | Cys        | lle        | Leu<br>810 | Asn        | Ser              | Phe        | Tyr        | Gly<br>815 | Tyr         |
| Val        | Met        | Arg        | Arg<br>820 | Gly        | Ala        | Arg        | Trp        | His<br>825 | Ser        | Met        | Pro              | Met        | Ala<br>830 | Gly        | He          |
| Val        | Cys        | Leu<br>835 | Thr        | Gly        | Ser        | Asn        | lle<br>840 | He         | Thr        | Lys        | Ala              | Arg<br>845 | Glu        | He         | He          |
| Glu        | Arg<br>850 | Val        | Gly        | Arg        | Pro        | Leu<br>855 | Glu        | Leu        | Asp        | Thr        | Asp<br>860       | Gly        | lie        | Trp        | Cys         |
| lle<br>865 | Leu        | Pro        | Gly        | Ser        | Phe<br>870 | Pro        | GIn        | Glu        | Phe        | Thr<br>875 | lle              | His        | Thr        | Ser        | His<br>880  |
| Glu        | Lys        | Lys        | Lys        | Lys<br>885 | lle        | Asn        | He         | Ser        | Tyr<br>890 | Pro        | Asn              | Ala        | Val        | Leu<br>895 | Asn         |
| Thr        | Met        | Val        | Lys<br>900 | Asp        | His        | Phe        | Thr        | Asn<br>905 | Asp        | GIn        | Tyr              | His        | Glu<br>910 | Leu        | Arg         |
| Lys        | Asp        | Lys<br>915 | Glu        | Asn        | Asn        | Leu        | Pro<br>920 | Lys        | Tyr        | Asp        | He               | Arg<br>925 | Asp        | Glu        | <b>A</b> sn |

| Ser          | 930          | rne         | rne         | Giu                  | vai         | 935         | шу           | rro         | ıyr         | Leu         | 940 <sub>.</sub> | wet.        | Vai         | LGU          | 710         |
|--------------|--------------|-------------|-------------|----------------------|-------------|-------------|--------------|-------------|-------------|-------------|------------------|-------------|-------------|--------------|-------------|
| Ala<br>945   | Ala          | Lys         | Glu         | Glu                  | Gly<br>950  | Lys         | Lys          | Leu         | Lys         | Lys<br>955  | Arg              | Tyr         | Ala         | Val          | Phe<br>960  |
| <b>A</b> sn. | Phe          | Asp         | Gly         | Thr<br>965           | Leu         | Ala         | Glu          | Leu         | Lys<br>970  | Gly         | Phe              | Glu         | Val         | Lys<br>975   | Arg         |
| Arg          | Gly          | Glu         | Leu<br>980  | Gin                  | Leu         | He          | Lys          | Asn<br>985  | Phe         | Gln         | Ser              | Ser         | Va I<br>990 | Phe          | Glu         |
| Ala          | Phe          | Leu<br>995  | Ala         | Gly                  | Ser         |             | Leu<br>1000  | Glu         | Glu         | Cys         |                  | Ala<br>1005 | Ser         | Val          | Ala         |
|              | Va I<br>1010 | Ala         | Asp         | Tyr                  |             | Leu<br>1015 | Asp          | Val         | Leu         |             | Ser<br>1020      | Arg         | Gly         | Ser          | <b>A</b> sn |
| Leu<br>102   |              | Asp         | Ser         |                      | Leu<br>1030 | Phe         | Glu          | Leu         |             | Ser<br>1035 | Glu              | Asn         | Lys         |              | Met<br>1040 |
| Ser          | Lys          | Lys         |             | Glu<br>1 <b>04</b> 5 | Glu         | Tyr         | Gly          |             | Gin<br>1050 | Lys         | Ser              | Thr         |             | l le<br>1055 | Ser         |
| Thr          | Ala          |             | Arg<br>1060 | Leu                  | Ala         | Glu         | Phe          | Leu<br>1065 | Gly         | Glu         | Gln              |             | Val<br>1070 | Lys          | Asp         |
| Ala          |              | Leu<br>1075 | Ala         | Cys                  | Lys         |             | l le<br>1080 | He          | Ser         | Lys         |                  | Pro<br>1085 | Glu         | Gly          | Ala         |
|              | Va1<br>1090  |             | Glu         | Arg                  |             | lle<br>1095 | Pro          | Leu         | Ala         |             | Phe<br>1100      | Gin         | Ser         | Glu          | Pro         |
| Ser<br>110   |              | Arg         | Arg         |                      | His<br>1110 | Leu         | Arg          | Arg         |             | Leu<br>1115 |                  | Asp         | Asn         |              | Met<br>1120 |

| Gly Asp         | Ala           |             | i le<br>1125 | Arg         | <b>A</b> sp | Val         |              | Asp<br>1130 | Trp         | Asn         | Tyr         |             | 1 le<br>1135 | Glu         |
|-----------------|---------------|-------------|--------------|-------------|-------------|-------------|--------------|-------------|-------------|-------------|-------------|-------------|--------------|-------------|
| Arg Lei         | _             | Gly<br>1140 | Thr          | He          | Gin         | =           | l le<br>1145 | lle         | Thr         | lle         |             | Ala<br>1150 | Ala          | Leu         |
| Gin Giy         | / Leu<br>1155 | Ala         | Asn          | Pro         |             | Pro<br>1160 | Arg          | Val         | Gln         |             | Pro<br>1165 | Asp         | Тгр          | Leu         |
| His Lys<br>1170 |               | Met         | Leu          |             | Lys<br>1175 | Asn         | Asp          | Val         |             | Lys<br>1180 | Gin         | Arg         | Arg          | lle         |
| Asn Glu<br>1185 | u Met         | Phe         |              | Ser<br>1190 | Arg         | Pro         | Lys          |             | Lys<br>1195 | Pro         | Leu         | Ala         |              | Glu<br>1200 |
| Giu Ası         | ) Lys         |             | Ala<br>1205  | Asp         | Met         | Glu         |              | Leu<br>1210 | Ala         | Gly         | Lys         |             | Gly<br>1215  | Gly         |
| Glu Gly         |               | Ala<br>1220 | Gly          | Cys         | Pro         |             | Va I<br>1225 | Thr         | Lys         | Arg         |             | Arg<br>1230 | lle          | Gln         |
| Leu Gli         | ı Glu<br>1235 |             | Asp          | Glu         |             | Glu<br>1240 | Ala          | Gln         | Pro         |             | Ala<br>1245 | Thr         | Thr          | Trp         |
| Arg Gli<br>1250 |               | Leu         | Gly          |             | Pro<br>1255 | Pro         | Pro          | lle         |             | Glu<br>1260 | Thr         | Arg         | Lys          | Thr         |
| l le Va<br>1265 | l Glu         | Trp         |              | Arg<br>1270 | Phe         | Gln         | Lys          |             | Lys<br>1275 | Trp         | Lys         | Trp         |              | GIn<br>1280 |
| Asp Gi          | n Arg         |             | Arg<br>1285  | Asn         | Arg         | Gln         |              | Ser<br>1290 | Lys         | Arg         | Thr         |             | Gly<br>1295  | Glu         |
| Asp Pro         |               | Tyr<br>1300 |              | Gly         | Arg         |             | Leu<br>1305  | Arg         | Arg         | Ala         |             | Arg<br>1310 | Thr          | Leu         |
| Leu As          | o Gin         | Pro         | Trp          | Gln         | He          | Val         | Gin          | Leu         | Val         | Pro         | Val         | Asp         | Asp          | Leu         |

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| 1315                  |                       | 1320                | 132                                 | 25                     |
|-----------------------|-----------------------|---------------------|-------------------------------------|------------------------|
| Gly His Phe 7<br>1330 |                       | Ala Leu IIe<br>335  | Gly Glu Glu <sub>.</sub> Le<br>1340 | eu His Lys IIe         |
| Lys Leu Thr \<br>1345 | Val Pro Arg<br>1350   | lle Phe Tyr         | Val Asn Gin Ar<br>1355              | rg Ser Ala Ala<br>1360 |
| Pro Pro Glu (         | Glu Gly Gln i<br>1365 |                     | Lys Val Asn Ar<br>370               | rg Val Leu Pro<br>1375 |
|                       | Pro Val Phe A<br>380  | Asn Leu Tyr<br>1385 | Arg Tyr Ser Va                      | al Pro Glu Gin<br>1390 |
| Leu Phe Arg 1<br>1395 | Asp Asn Ser I         | Leu Gly Met<br>1400 | Leu Ala Asp Le<br>140               | eu Ala Thr Pro<br>05   |
| Asp lie Glu (<br>1410 |                       | Glu Thr Gln<br>415  | Met Thr Leu G<br>1420               | u Phe Arg Ala          |
| Leu Met Asp!<br>1425  | Met Gly Cys<br>1430   | lle Cys Gly         | Val Gin Arg Gi<br>1435              | u Glu Ala Arg<br>1440  |
| Arg Leu Ala (         | Gin Leu Ala<br>1445   |                     | Leu Glu Thr Pt<br>450               | ne Ser lie Glu<br>1455 |
|                       | Gin Gly Pro /<br>460  | Arg Leu Arg<br>1465 | Ser Asn lle Tr                      | rp Leu Ala Pro<br>1470 |
| Thr lle Asp (<br>1475 | Cys Ala Lys           | Ser Thr Cys<br>1480 | lie Ser ile Tr<br>148               | nr His Arg Arg<br>85   |
| Pro Arg Arg A         |                       | Ser Leu lle<br>495  | Pro Met Pro Se<br>1500              | er Lys Lys Ala         |
| Phe Val Phe A         | Ala Leu Asp<br>1510   | Thr Val Arg         | Ala Asn Gin Me<br>1515              | et Pro Asn Met<br>1520 |

| Arg         | Gin        | Leu         | _           | Thr<br>1525 | Ala         | Glu                  | Arg                 |             | Ala<br>1530 | Leu         | Leu         | Lys                  |             | Leu<br>1535 | Thr         |
|-------------|------------|-------------|-------------|-------------|-------------|----------------------|---------------------|-------------|-------------|-------------|-------------|----------------------|-------------|-------------|-------------|
| Ala         | Glu        |             | Gln<br>1540 | Asp         | Lys         | He                   |                     | Val<br>1545 | Glu         | Asp         | Tyr         |                      | Phe<br>1550 | Glu         | Val         |
| Leu         |            | Glu<br>1555 | Val         | Asp         | Val         |                      | GIn<br>1560         | He          | Tyr         | Arg         |             | lle<br>1565          | Gln         | Arg         | Ala         |
|             | Thr<br>570 | Thr         | Tyr         | Lys         |             | Glu<br>1 <b>5</b> 75 | His                 | Gln         | Gly         |             | Pro<br>1580 | Thr                  | lle         | Leu         | Cys         |
| Leu<br>1585 |            | Thr         | Ala         | Leu         | Ser<br>1590 | Ala                  | Arg                 | Lys         |             | Ser<br>1595 | Leu         | Ala                  | Met         |             | lle<br>1600 |
| Leu         | Leu        | Glu         |             | Pro<br>1605 | Gin         | Ala                  | Gln                 |             | His<br>1610 | lle         | Ser         | Asp                  |             | Ala<br>1615 | Ser         |
| Leu         | Leu        |             | Gly<br>1620 | Leu         | Asp         | Trp                  |                     | Arg<br>1625 | Gln         | Gly         | Ser         |                      | Ala<br>1630 | Val         | He          |
| Arg         |            | Phe<br>1635 | Leu         | Asn         | Leu         |                      | <b>A</b> sn<br>1640 | Vai         | Leu         | Asp         |             | Met<br>1 <b>64</b> 5 | Leu         | Asp         | Gln         |
|             | Arg<br>650 | Tyr         | Phe         | His         |             | Pro<br>1 <b>65</b> 5 | He                  | Gly         | Asn         |             | Pro<br>1660 | Pro                  | Asp         | Thr         | Val         |
| Leu<br>1665 |            | Gly         | Ala         | Asp         | Leu<br>1670 | Phe                  | Phe                 | Ala         | _           | Leu<br>1675 | Leu         | Gin                  | Arg         |             | Asn<br>1680 |
| Phe         | Val        | Leu         |             | Trp<br>1685 | Ser         | Ala                  | Ser                 |             | Arg<br>1690 | Pro         | Asp         | Leu                  |             | Gly<br>1695 | Arg         |
| Glu         | Ala        |             | Asp<br>1700 | Ser         | Arg         | Leu                  |                     | Ala<br>1705 | Glu         | Phe         | Glu         |                      | Ser<br>1710 | He          | Ser         |

| <b>Val Val G</b> in<br>1715 |                     | Gly Phe 1<br>1720        | Tyr Pro Asp Va         | l Cys Val Glu Leu<br>1725 |
|-----------------------------|---------------------|--------------------------|------------------------|---------------------------|
| Ala Leu Asp<br>1730         |                     | Val Ser <i>I</i><br>1735 | Ala Leu Leu Gl<br>174  | n Ser Thr Arg lle<br>O    |
| Gin Giu Met<br>1745         | Glu Gly Ala<br>1750 | Ser Ser /                | Ala lie Thr Ph<br>1755 | e Asp Val Met Pro<br>1760 |
| Gin Val Ser                 | Leu Glu Glu<br>1765 | Met lle (                | Gly Thr Val Pr<br>1770 | o Ala Ala Thr Leu<br>1775 |
|                             | Asp Glu Thr<br>1780 |                          | Cys Ser Ala Al<br>785  | a Phe Arg Val Met<br>1790 |
| Arg Ser Met<br>1795         |                     | Trp Leu /                | Arg Glu Val Se         | r lle Asn Arg Asr<br>1805 |
| lle Phe Ser                 | Asp Phe Gin         | lie Val I                | lis Phe Tyr <b>A</b> r | g Trp Val Arg Ser         |
| 1810                        |                     | 1815                     | 182                    | 0                         |
| Ser Asn Ala<br>1825         | Leu Leu Tyr<br>1830 | Asp Pro /                | Ala Leu Arg Ar<br>1835 | g Ser Leu Asn Asr<br>1840 |
| <b>Leu Met Arg</b>          | Lys Met Phe<br>1845 | Leu Arg                  | lle lle Ala Gl<br>1850 | u Phe Lys Arg Leu<br>1855 |
|                             | lle lle Tyr<br>1860 |                          | Phe Asn Arg II<br>365  | e lle Leu Ser Ser<br>1870 |
| Gly Lys Lys<br>1875         |                     | Asp Ala l<br>1880        | ∟eu Gly Tyr Va         | l Asp Tyr lle Val<br>1885 |
| Gln Ser Leu<br>1890         |                     | Glu Met F<br>1895        | Phe His Ser II<br>190  | e Gin Leu Ser Phe<br>O    |

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| Glu Gln<br>1905 | Cys         | Trp         |             | Phe<br>1910 | Met         | Leu         | Trp         |             | Asp<br>1915 |             | Ala         | Asn         |             | Ser<br>1920  |
|-----------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|--------------|
| Gly lle         | Arg         |             | Lys<br>1925 | Leu         | Pro         | Lys         |             | lle<br>1930 | Asp         | Glu         | Thr         |             | Ser<br>1935 | Ser          |
| lle Val         |             | Thr<br>1940 | Thr         | Met         | lle         |             | Asp<br>1945 | Ser         | Glu         | Arg         |             | GIn<br>1950 | Asp         | Asp          |
| Asp Glu         | Asp<br>1955 |             | Glu         | Glu         |             | Ser<br>1960 | Glu         | Asn         | Arg         |             | Pro<br>1965 | Val         | Glu         | Ser          |
| Asn Glu<br>1970 |             | Glu         | Gln         |             | GIn<br>1975 | Glu         | Asp         | Glu         |             | Ser<br>1980 | Leu         | Glu         | Leu         | Asn          |
| Trp Thr<br>1985 | He          | Gly         |             | His<br>1990 | Leu         | Pro         | Asp         |             | Asn<br>1995 | Glu         | Cys         | Arg         |             | Lys<br>2000  |
| Phe Glu         | Ser         |             | Leu<br>2005 | Thr         | Leu         | Phe         |             | GIn<br>2010 | Ser         | Leu         | Ala         |             | Lys<br>2015 | Lys          |
| Thr Thr         |             | Gin<br>2020 | Ala         | lle         | Lys         |             | lle<br>2025 | Ser         | His         | Cys         |             | Phe<br>2030 | Asp         | Phe          |
| lle Leu         | Lys<br>2035 | Leu         | His         | Lys         |             | Tyr<br>2040 | Gly         | Lys         | Gly         |             | Pro<br>2045 | Ser         | Pro         | Gly          |
| Leu Glu<br>2050 | Leu         | He          | Arg         |             | Leu<br>2055 | lle         | Lys         | Ala         |             | Ser<br>2060 | Val         | Asp         | Lys         | Thr          |
| Leu Ala<br>2065 | Glu         | Gln         |             | Asn<br>2070 | Glu         | Leu         | Arg         |             | Asn<br>2075 | Met         | Leu         | Arg         |             | Va I<br>2080 |
| Gly lle         | Gly         |             | Phe<br>2085 | Ser         | Asp         | Leu         |             | G1u<br>2090 | Trp         | Glu         | Asp         |             | Cys<br>2095 | Asp          |
| Ser His         | He          | He          | Asn         | Glu         | Val         | He          | Cys         | Lys         | Ala         | Cys         | Asn         | His         | Cys         | Arg          |

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2100 2105 2110

Asp Leu Asp Leu Cys Lys Asp Lys His Arg Ala Met Lys Asp Gly Cys 2115 2120 2125

Tyr Val Ala Tyr Asp Asn Glu Glu Ile Glu Met Arg Met Leu Asp Ala 2130 2135 2140

Leu Gin Arg Lys Met Met Ser Tyr Val Leu Gin Asp Leu Arg Cys Ser 2145 2150 2155 2160

Arg Cys Ser Glu IIe Lys Arg Glu Asn Leu Ala Glu Phe Cys Thr Cys 2165 2170 2175

Ala Gly Asn Phe Val Pro Leu IIe Ser Gly Lys Asp IIe Gln Thr Leu 2180 · 2185 2190

Leu Gly Thr Phe Asn Lys Val Ala Ala Asn His Lys Met Gln Leu Leu 2195 2200 2205

Gin Gin Thr Vai His Gin Ala Leu Thr Thr Pro Arg 2210 2215 2220

<210> 67

<211> 26

<212> DNA

<213> Artificial Sequence

<220>

<223> primer

**<400> 67** 

gactagtggc tatcttgtgg cgggaa

26

<210> 68

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|                    |   |     |
| <220>              |   |     |
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|                    |   |     |
| <b>&lt;400&gt;</b> |   |     |
| ggaatt             | cctt gtcccgtgtc aggtca                                      | 26  |
|                    |   |     |
| <210>              | 60  |     |
| ⟨211⟩              |   |     |
| <212>              |   |     |
|                    | Artificial Sequence   |     |
|                    |   |     |
| <220>              |   |     |
| <223>              | primer  |     |
|                    |   |     |
| <b>&lt;400&gt;</b> | 69  |     |
| cagaac             | tttg ccctcccata cctc  | 24  |
|                    |   |     |
| (2.4.2)            |   |     |
| <210>              |   |     |
| <211><br><212>     |   |     |
|                    | Mus musculus  |     |
| VZ13/              | nus nuscutus  |     |
| <b>&lt;400&gt;</b> | 70  |     |
|                    | tgta agoggogaca aggaccaggo cotggggtgo coccaaagog ggotogaggg | 60  |
|                    |   |     |
| cacctc             | tggg atgaggacga gccttcgccg tcgcagtttg aggcgaacct ggcactgctg | 120 |
|                    |   |     |
| gaggaa             | atag aggotgagaa ocggotgoag gaggoagagg aggagotgoa gotgococca | 180 |
|                    |   |     |
| gagggc             | accg tgggtgggca gttttccact gcagacattg accctcggtg gcggcggccc | 240 |

| 300  | ggagattgac | tocagcagct | ccctcatct  | cagcacggag | ccctggaccc | accetacgtg |
|------|------------|------------|------------|------------|------------|------------|
| 360  | gaactcagtg | tgccatcccg | gaagggcccc | acccctgcca | gctcagcacc | cactatgtgg |
| 420  | ccacatacag | ccgtctgctg | gaaggettet | ggtcaccgat | gggcctttgg | cccatactga |
| 480  | gcacctgagt | ttggggccga | cctcctggtt | caccccgcg  | cctacttcta | ggctttgccc |
| 540  | gaaggagctc | agcgcggtgg | agccgggacc | cgcagccatc | aggagctgaa | gagctgcagc |
| 600  | tgggtaccac | agagcatgtt | tgctcccgtg | aatagagota | cagtgctggc | tcagggccgg |
| 660  | tatggcacca | tacccegcct | accetggeac | tctccgcatc | cttctccatt | ggtcatggcc |
| 720  | gagcttcgca | tgggcacccc | gtgccaggcc | gggtgtccga | ttctggaaca | gcccgccgcc |
| 780  | cattgtggga | tggatgctga | cggttcatgg | ctttgagatc | ccaacgtgga | ccctacgaag |
| 840  | gaaggccacc | gggcggagaa | tacgttcgga | agctggaaag | tggagctgcc | tgcaactggt |
| 900  | accggagggg | tcagtcaccc | tcagatgtga | cgtgctgtgg | tggaggtgga | ctgtgtcagc |
| 960  | tgctggccga | acatogagtg | cttagcttcg | cctgcgtgtg | gcattgcacc | cagtggcagc |
| 1020 | totggggctg | agatotgtto | cccgtgatcc | tgagcgtgac | tccctgagcc | aaaggcatct |
| 1080 | ctgtgccccc | cgctgcggcc | ctggcactca | attettgegt | agccggagcc | cgctgggggg |
| 1140 | ggcctgggcc | acctgctcca | cgggaagaag | gagctatgag | ccaaagtgca | atcctgggtg |
| 1200 | gaactttgcc | acaacattca | atcaccggct | ccctgacgtg | ttgccatgga | gacttcatcc |
| 1260 | tttcctgggc | accgottccc | ctaaaggtgg | ggcacaggcc | tcatctctcg | ctcccatacc |
| 1320 | gcaggtcggc | tccaatcaag | gactcctcct | caacatcogt | gtctccgctc | cgcgtgactg |
| 1380 | gctgcaggtg | agatggatat | ggtcgcgttc | cagcatggtg | gtaaggtgat | cggcgggaca |

1440 ctgcttcggg aacacaagct ccgctcctac acgctcaacg ctgtgagttt ccacttcctg 1500 ggcgagcaga aggaggacgt tcagcacagc atcatcaccg acctgcagaa tgggaacgaa cagacgegec geogectgge egtgtactge etgaaggaeg cetttetgec acteegaeta 1560 1620 ctagagcgcc ttatggtgct ggtgaataat gtggagatgg cgcgtgtcac gggtgtaccc 1680 cttgggtacc tgctcacccg gggccagcag gtcaaggtcg tgtctcagct gctgcgccag gccatgcgcc aggggctgct gatgcctgtg gtgaagaccg agggcagtga ggactacacg 1740 1800 ggagccacag tcattgagcc cctcaaaggg tactatgacg tccccattgc caccctggac 1860 ttctcctcct tgtacccatc catcatgatg gcccataatc tgtgctacac cacgctgctc 1920 cgacctgggg ctgcccagaa gctgggcctt aaaccagatg agttcatcaa gacacccact ggggatgagt ttgtgaagtc atctgtacgg aagggcctcc tgccccagat cctggagaat 1980 2040 ctgctgagtg cccgcaagag ggccaaggct gagctggctc aggagacgga ccccctgcgg 2100 cgacaggtct tggacggccg gcaactggca ctaaaagtga gtgccaactc cgtatatggc 2160 ttcactggtg cccaggtggg caagctgcca tgtttggaga tctcccagag tgtcactggg ttcgggcggc agatgattga gaaaaccaag cagcttgtgg agtccaagta caccgtggaa 2220 2280 aatggctacg atgccaacgc caaggtagtc tacggtgaca cggactctgt gatgtgccgg 2340 tttggcgtct cctctgtggc tgaagcaatg tctctggggc gggaggctgc aaactgggta tocagteact teccateace cateeggetg gagttegaga aggtttactt eccatacetg 2400 ctcatcagca agaagegeta tgctggcctg ctcttctcct cccgctctga tgcccatgac

| aaaatggact gcaagggcct | ggaggctgtg   | cgcagggaca | actgtcccct | ggtggccaac | 2520 |
|-----------------------|--------------|------------|------------|------------|------|
| ctcgttacat cctctctgcg | ccggatcctc   | gtggaccggg | accetgatgg | ggcagtagcc | 2580 |
| catgccaagg acgtcatctc | ggacctgctg   | tgcaaccgca | tagacatoto | ccagctggtc | 2640 |
| atcaccaaag agttgacccg | cgcagcagca   | gactatgctg | gcaagcaggc | tcacgtggag | 2700 |
|                       |              |            |            |            |      |
| ctggctgaga ggatgaggaa | gegegaeeee   | ggcagtgcgc | ccagcctggg | tgaccgagtc | 2760 |
| ccctatgtga tcattggtgo | tgctaagggt   | gtggccgcct | acatgaagtc | ggaggacccc | 2820 |
| ctgtttgtgc tggagcacag | cctgcccatc   | gacactcagt | actacctgga | gcagcagctg | 2880 |
| gccaagccgc tcttgcgcat | ctttgagccc   | atcctgggtg | agggccgtgc | agagtctgtg | 2940 |
| ctgctgcgcg gtgaccacac | acgatgcaag   | actgtgctca | ccagcaaggt | gggcggcctc | 3000 |
| ttggccttca ccaagcgccg | caactgttgc   | attggctgcc | gctccgtaat | cgaccatcaa | 3060 |
| ggagccgtgt gtaagttctg | tcagccacgg   | gagtcggagc | tctatcagaa | ggaggtgtca | 3120 |
| cacctgaatg ccttggaaga | acggttctct   | cgcctctgga | cacagtgtca | acgctgccag | 3180 |
| ggcagcttgc atgaggacg  | : catctgtacc | agccgtgact | gtcccatctt | ctacatgcgc | 3240 |
| oagaaggtgc gcaaggacct | ggaagaccag   | gaacggctgc | tgcagcgctt | tggaccgccc | 3300 |
| ggccctgagg cctggtga   |              |            |            |            | 3318 |

<210> 71

<211> 29

<212> DNA

<213> Artificial Sequence

| <b>&lt;220&gt;</b> |                           |    |
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| <223>              | primer                    |    |
|                    |                           |    |
| <b>&lt;400&gt;</b> | 71                        |    |
| tccccg             | cggc tgcagaggat tttccacag | 29 |
|                    |                           |    |
|                    |                           |    |
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| <211>              |                           |    |
| <b>&lt;212&gt;</b> |                           |    |
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|                    | ·                         |    |
| <220>              |                           |    |
| <b>&lt;223&gt;</b> | primer                    |    |
|                    |                           |    |
| <b>&lt;400&gt;</b> |                           |    |
| ggacta             | gtat ggtatgcaca acagcctc  | 28 |
|                    |                           |    |
| <210>              | פד                        |    |
| <211>              |                           |    |
| <212>              |                           |    |
|                    | Artificial Sequence       |    |
| W2107              | 78 CTTOTAL COGGOTOC       |    |
| ⟨220⟩              |                           |    |
| ⟨223⟩              | primer                    |    |
|                    |                           |    |
|                    |                           |    |
| <b>&lt;400&gt;</b> | 73                        |    |
| tocccg             | cgga gtggttgtgg gagacttac | 29 |
|                    |                           |    |
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| <210>              | 74                        |    |
| <211>              | 28                        |    |
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<220>

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|--------------------|--------------------------|----|
| <223>              | primer                   |    |
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| <b>&lt;400&gt;</b> | 74                       |    |
| ggacta             | gtca gtcccacagt cccaaagt | 28 |
|                    |                          |    |
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| <210>              | 75                       |    |
| <211>              | 18                       |    |
| <212>              | DNA                      |    |
| <213>              | Artificial Sequence      |    |
|                    |                          | •  |
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| <223>              | primer                   |    |
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| <400>              | 75                       |    |
| ctgaga             | gtga tgaggttc            | 18 |
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|                    |                          |    |
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| <223>              | primer                   |    |
| (100)              |                          |    |
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| ctaatc             | gcca tcttccagca g        | 21 |
|                    | ·                        |    |
| <210>              | 77                       |    |
| <211>              | 20                       |    |
| <212>              |                          |    |
|                    | Artificial Sequence      |    |
| V213/              | A LITTOTAL SEQUENCE      |    |
|                    |                          |    |

| <b>&lt;223&gt;</b> | primer              |    |
|--------------------|---------------------|----|
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|                    |                     |    |
| <210>              | 78                  |    |
| <211>              | 20                  |    |
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| <213>              | Artificial Sequence |    |
| (000)              |                     |    |
| ⟨220⟩              |                     |    |
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|                    |                     |    |
| <210>              | 79                  |    |
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| ⟨212⟩              |                     |    |
| ⟨213⟩              | Artificial Sequence |    |
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| <220>              | •                   |    |
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| go cogo            |                     |    |
|                    |                     |    |
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| <211>              | 20                  |    |
| <b>&lt;212&gt;</b> |                     |    |
| <213>              | Artificial Sequence |    |
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|                    | primer              |    |
|                    |                     |    |

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**<400> 80** 

ccaacgctat gtcctgatag

20

<210> 81

<211> 5725

<212> DNA

<213> Mus musculus

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ccccggcata gtctcacaag agacagctac atctgggtcc tttcgataaa atcttgctag 840 900 tgtatgcaat ggtgtcagcg tttggatgct gattatgggg tggatccctg gataaggcag 960 tototacatg gtocatoctt toatotoago tocaaacttt gtototgtaa ctoottocaa 1020 gggtgttttg ttcccactc taaggagggg catagtgtcc acacttcagt cttcttttt 1080 catgagtttc atgtgtttag gaaattgtat cttatatctt gggtatccta ggttttgggc taatatocac ttatcagoga gtacatattg tgtgagttcc tttgtgaatg tgttacctca 1140 ctcaggaaga tgccctccag gtccatccat ttggctagga atttcataaa ttcattcttt 1200 1260 ttaatagctg agtagtactc cattgtgtag atgtaccaca ttttctgtat ccattcctct gttgaggggc atctgggttc tttccagctt ctggctatta taaataaggc tgctatgaac 1320 1380 atagtggagc atgtgtcctt cttaccagtt ggggcatctc ctggatatat gcccaggaga ggtattcctg gatcctccgg tagtactatg tccaattttc taaggaaccg ccagatggat 1440 1500 ttccagagtg gttgtacaag cctgcaatcc caccaacaat ggaggagtgt tcctctttct 1560 ccacatcctc gccagcatct gctgtcacct gaatttttga tcttagccat tctgactggt 1620 gtgaggtgga atctcagggt tgttttgatt tgcatttccc tgatgattaa ggatgttgaa 1680 cattttttca ggtgcttctc tgccattcgg tattcctcag gtgagaattc tttgttcagt 1740 tetgagecce attittitaa tggggttatt tgattitetg aagtecacet tettgagtte 1800 tttatatatg ttggatatta gtcccctatc tgatttagga taggtaaaga tcctttccca atotgttggt ggtctctttg tcttattgac agtgtctttt gccttgcaga aactttggag 1860 1920 

| caggaatttt | tcccctgtgc | ccatatcttc | aaggcttttc | cccactttct | cctctataag | 1980 |
|------------|------------|------------|------------|------------|------------|------|
| tttcagtgtc | totggtttta | tgtgaagttc | tttgatccat | ttagatttga | ccttagtaca | 2040 |
| aggagataag | tatggatcga | ttcgcattct | tctacatgat | aacaaccagt | tgtgccagca | 2100 |
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| cctaggtttt | ttgttattcc | agatgaattt | gcaaattgct | ccttctaatt | cgttgaagaa | 2400 |
| ttgagttgga | attttgatgg | ggattgcatt | gaatctgtag | attgcttttg | gcaagatagc | 2460 |
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| atcttcttta | atttctttct | tcagagactt | gaagtttta  | tcatacatat | ctttcacttc | 2580 |
| ctcagttaga | gtcacgccga | gatattitat | attatttgtg | actattgaga | agggtgttgt | 2640 |
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| tgagttaatt | ttatatccag | ctacttcacc | aaagctgttt | atcaggttta | ggagttctct | 2760 |
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| gacttcctct | tttccaattt | gtatcccctt | gatctccttt | tgttgtcgaa | ttgctcagca | 2880 |
| ctgcagtatt | cttaacaaag | ttgattctca | ttagaaaata | aagctgccat | atgaccatct | 2940 |
| aacacttagt | aactgggaaa | atttgatttg | atctggtgat | ttgtcacago | aataacaaag | 3000 |
|            |            |            |            | _          |            |      |

| taactatcac | aatggctgaa | acaagtotgo | tcacttggaa | tcaacacttt | ttgtcccagg | 3060 |
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| agcaacagag | tottcagtat | caggggccta | aatttagaga | tgaaggtttg | gacctgtagc | 3120 |
| ctgaacctct | tgcagaagtg | atgtattaaa | aggctgatca | acaatggtcg | ctacaagagt | 3180 |
| actccaggcc | agccaatcca | acaageccaa | taccttgaag | gatgaaaccc | atgcccaaga | 3240 |
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| ctagatgcta | gtgttcctct | aagcatgaat | aactgatata | tattcatgct | gctctacctc | 3360 |
| agatattcag | acttttcccc | ctggttttca | gtaaatcaag | ttggatcact | ccactcttct | 3420 |
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| caggtgccaa | gtggttccaa | atagctgggt | gcagtttgtc | tcaggggcac | ctctagaggg | 3780 |
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| gggcgcagtg | tacatgggtg | ttctccagta | acagtcttgg | ctgggaagga | gagctagatt | 3960 |
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| aacaaatagt | aaagctctgt | tgttgttgtt | gataagtagc | aaattaattt | agggggagca | 4080 |

| gtttgggggg ctgatgggtc | ttgttctgcc | attcaggctg | ggctgaagct | cctagggtct | 4140 |
|-----------------------|------------|------------|------------|------------|------|
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| ctaggettee aateaggttt | cacattttgc | ttctagacag | aaatataatt | tectgaaacc | 4260 |
| tttcgttttg aaatgatgtc | attattgggc | cccgcactc  | attgctgctg | acgccccct  | 4320 |
| cccactatcc ctcgggctga | aatgtcctga | ttgggtgtca | acgctcattt | gcactgatgc | 4380 |
| tccccctta atccctggag  | ctgaaatgtc | ctgattgggt | gccaaaattt | tttccactga | 4440 |
| tgcctctccc atcccattag | cactaggact | gaaatgtcat | gattgggcgc | aaaaactaat | 4500 |
| ttccaattat gcccggcccc | tccgattatc | cctggtgctg | agatgtcatg | attggggcca | 4560 |
| acactcattt ctgctgatgg | ccctcctctc | ccattagacc | tggacctgaa | tccgtgtcat | 4620 |
| gattggacgc caacactcat | ttccactgat | gcccggccc  | tcccattagt | cctgaggctg | 4680 |
| aacccctgtc atgattgggc | tccaatcctc | atttcaggtg | atgotoagot | cctcccatca | 4740 |
| gccctgtggc tgaatctctg | tcacgattgg | gcccaaccc  | tcatttctgc | tgaggcccag | 4800 |
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| gctgatgccc cgcccatgcc | attacccttg | ggcctaaatc | cctgtcatga | ttgaactcca | 4920 |
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| togggotgat cootgtoatg | attgggcccc | aaccctcatt | toggotgagg | cccactcct  | 5160 |
| ccaattagcc ctgaggctga | atccatgtca | tgagtaggca | ccaactctca | ttaacactga | 5220 |

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| ggcaccaaac cgcaattcca ctaaagcccc acccctcca ttaccatcct gccgaaacca stgtcttgtca tgattgggcc ccaaccctcc tttccactga tgccccccc tcccttaagc scctcctgctg agaccacatc ttgattgggc actaacactc atttccgctg atgcccacca sccccatttg ccctgggact taaaccctgt cgtgattggg tgtcaaccct catttccggt gatgccccgc ctcttcctat aaatcctggc gctcaaatac tgtggtcggt gggcaggaac sagccatttgg atcactgcct gcagcctagc ggttgagctg ctctggcgat catctgttct sagccatttgg atcactgcct gcagcctagc ggttgagctg ctctggcgat catctgttct | 280 |
|---|-----|
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| cetectgetg agaceacate ttgattggge actaacacte attteegetg atgeceacea geteceatttg ecetgggact taaaceetgt egtgattggg tgteaaceet cattteeggt gatgeceege etetteetat aaateetgge geteaaatae tgtggteggt gggeaggaac gagecatttgg ateaetgeet geageetage ggttgagetg etetggegat eatetgttet gaggtaettt gggactgtgg gactg   | 400 |
| ctcccatttg ccctgggact taaaccctgt cgtgattggg tgtcaaccct catttccggt gatgccccgc ctcttcctat aaatcctggc gctcaaatac tgtggtcggt gggcaggaac gaccatttgg atcactgcct gcagcctagc ggttgagctg ctctggcgat catctgttct gaggtacttt gggactgtgg gactg   | 460 |
| gatgececge etetteetat aaateetgge geteaaatae tgtggteggt gggeaggaac gageeatttgg ateaetgeet geageetage ggttgagetg etetggegat eatetgttet gaggtaettt gggaetgtgg gaetg  | 520 |
| agccattigg atcactgcct gcagcctagc ggttgagctg ctctggcgat catcigtict { gaggtacttt gggactgigg gactg {   | 580 |
| gaggtacttt gggactgtgg gactg   | 640 |
| 3.00 - 0.00 3 0.00  | 700 |
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| \Z10/ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \   |     |
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**<400> 82** 

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30

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| agcaac                         | gago tottatgatt ggtttatotg 30                                   |
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| <223>                          | primer  |
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| atttgo                         | tgtc gataatatca gatttcttgg 30                                   |
|                                |   |
|                                |   |
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| <211>                          | 20  |
| (010)                          | 20  |
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|                                |   |
|                                | DNA   |
|                                | DNA   |
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| <220≻  |     |  |  |  |  |  |  |  |  |  |  |  |
|--|-----|--|--|--|--|--|--|--|--|--|--|--|
| <221> CDS  |     |  |  |  |  |  |  |  |  |  |  |  |
| <b>⟨222⟩ (24) (3341)</b>   |     |  |  |  |  |  |  |  |  |  |  |  |
| <400> 86   |     |  |  |  |  |  |  |  |  |  |  |  |
| ggcgggaaaa gctgtttgag gcg atg gat tgt aag cgg cga caa gga cca ggc 53 |     |  |  |  |  |  |  |  |  |  |  |  |
| Met Asp Cys Lys Arg Arg Gin Giy Pro Giy                              |     |  |  |  |  |  |  |  |  |  |  |  |
| 1 5 10   |     |  |  |  |  |  |  |  |  |  |  |  |
| cct ggg gtg ccc cca aag cgg gct cga ggg cac ctc tgg gat gag gac      | 101 |  |  |  |  |  |  |  |  |  |  |  |
| Pro Gly Val Pro Pro Lys Arg Ala Arg Gly His Leu Trp Asp Glu Asp      |     |  |  |  |  |  |  |  |  |  |  |  |
| 15 20 25   |     |  |  |  |  |  |  |  |  |  |  |  |
| gag cct tcg ccg tcg cag ttt gag gcg aac ctg gca ctg ctg gag gaa      | 149 |  |  |  |  |  |  |  |  |  |  |  |
| Giu Pro Ser Pro Ser Gin Phe Giu Ala Asn Leu Ala Leu Leu Giu Giu      |     |  |  |  |  |  |  |  |  |  |  |  |
| 30 35 40   |     |  |  |  |  |  |  |  |  |  |  |  |
| •  |     |  |  |  |  |  |  |  |  |  |  |  |
| ata gag gct gag aac cgg ctg cag gag gca gag gag gag ctg cag ctg      | 197 |  |  |  |  |  |  |  |  |  |  |  |
| lle Glu Ala Glu Asn Arg Leu Gin Glu Ala Glu Glu Glu Leu Gin Leu      |     |  |  |  |  |  |  |  |  |  |  |  |
| 45 50 55   |     |  |  |  |  |  |  |  |  |  |  |  |
| ccc cca gag ggc acc gtg ggt ggg cag ttt tcc act gca gac att gac      | 245 |  |  |  |  |  |  |  |  |  |  |  |
| Pro Pro Glu Gly Thr Val Gly Gly Gln Phe Ser Thr Ala Asp Ile Asp      |     |  |  |  |  |  |  |  |  |  |  |  |
| 60 65 70   |     |  |  |  |  |  |  |  |  |  |  |  |
|  |     |  |  |  |  |  |  |  |  |  |  |  |
| cct cgg tgg cgg cgc ccc acc cta cgt gcc ctg gac ccc agc acg gag      | 293 |  |  |  |  |  |  |  |  |  |  |  |
| Pro Arg Trp Arg Arg Pro Thr Leu Arg Ala Leu Asp Pro Ser Thr Glu      |     |  |  |  |  |  |  |  |  |  |  |  |
| 75 80 85 90  |     |  |  |  |  |  |  |  |  |  |  |  |
| _  |     |  |  |  |  |  |  |  |  |  |  |  |
| ccc ctc atc ttc cag cag ctg gag att gac cac tat gtg ggc tca gca      | 341 |  |  |  |  |  |  |  |  |  |  |  |
| Pro Leu IIe Phe Gin Gin Leu Glu IIe Asp His Tyr Val Gly Ser Ala      |     |  |  |  |  |  |  |  |  |  |  |  |
| 95 100 105   |     |  |  |  |  |  |  |  |  |  |  |  |
| cca ccc ctg cca gaa ggg ccc ctg cca tcc cgg aac tca gtg ccc ata      | 389 |  |  |  |  |  |  |  |  |  |  |  |

Pro Pro Leu Pro Giu Giy Pro Leu Pro Ser Arg Asn Ser Val Pro Ile

115

120

110

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| ctg        | agg         | gcc    | ttt  | ggg   | gtc | acc | gat | gaa   | ggc | ttc | tcc   | gtc | tgc | tgc | cac | 437 |
|------------|-------------|--------|------|-------|-----|-----|-----|-------|-----|-----|-------|-----|-----|-----|-----|-----|
| Leu        | Arg         | Ala    | Phe  | Gly   | Val | Thr | Asp | Glu   | Gly | Phe | Ser   | Val | Cys | Cys | His |     |
|            |             | 125    |      |       |     |     | 130 |       |     |     |       | 135 |     |     |     |     |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     |     |     |
| ata        | cag         | ggc    | ttt  | gcc   | CCC | tac | ttc | tac   | acc | CCC | gcg   | cct | cct | ggt | ttt | 485 |
| He         | Gln         | Gly    | Phe  | Ala   | Pro | Tyr | Phe | Tyr   | Thr | Pro | Ala   | Pro | Pro | Gly | Phe |     |
|            | 140         |        |      |       |     | 145 |     |       |     |     | 150   |     |     |     |     |     |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     |     |     |
| ggg        | gcc         | gag    | cac  | ctg   | agt | gag | ctg | cag   | cag | gag | ctg   | aac | gca | gcc | atc | 533 |
| Gly        | Ala         | Glu    | His  | Leu   | Ser | Glu | Leu | Gln   | Gin | Glu | Leu   | Asn | Ala | Ala | He  |     |
| 155        |             |        |      |       | 160 |     |     |       |     | 165 |       |     |     |     | 170 | •   |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     |     |     |
| agc        | cgg         | gac    | cag  | cgc   | ggt | ggg | aag | gag   | ctc | tca | ggg   | ccg | gca | gtg | ctg | 581 |
|            |             |        |      |       | Gly |     |     |       |     |     |       |     |     |     |     |     |
|            |             |        |      | 175   |     |     |     |       | 180 |     | •     |     |     | 185 |     |     |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     |     |     |
| gca        | ata         | gag    | cta  | tgc   | tcc | cgt | gag | agc   | atg | ttt | ggg   | tac | cac | ggt | cat | 629 |
|            |             |        |      |       | Ser |     |     |       |     |     |       |     |     |     |     |     |
|            |             |        | 190  |       |     |     |     | 195   |     |     |       |     | 200 |     |     |     |
|            |             |        |      |       |     |     |     |       | ٠   |     |       |     |     |     |     |     |
| ggc        | cct         | tct    | cca  | ttt   | ctc | cgc | atc | acc   | ctg | gca | cta   | ccc | cgc | ctt | atg | 677 |
|            |             |        |      |       | Leu |     |     |       |     |     |       |     |     |     |     |     |
| •          |             | 205    |      | _     |     |     | 210 |       |     |     |       | 215 |     |     |     |     |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     |     |     |
| gca        | cca         | gcc    | cgc  | CEC   | ctt | ctg | gaa | cag   | ggt | gtc | cga   | gtg | cca | ggc | ctg | 725 |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     | Leu |     |
|            | 220         |        |      | 0     |     | 225 |     |       | -   |     | 230   |     |     |     |     |     |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     |     |     |
| aac        | acc         | CCE    | ago  | ttc   | gca | CCC | tac | gaa   | gcc | aac | gtg   | gac | ttt | gag | atc | 773 |
|            |             |        |      |       |     |     |     |       |     |     | -     |     |     |     | Пe  |     |
| 235        |             |        | 00.  | , ,,, | 240 |     |     |       |     | 245 |       |     |     |     | 250 |     |
| <u>د</u> ن |             |        |      |       | 270 |     |     |       |     | 5   |       |     |     |     |     |     |
| can        | ++^         | ata    | orto | gat   | act | gar | att | o ta  | pps | tec | aac   | tge | ttø | gap | ctg | 821 |
|            |             |        |      |       |     |     |     |       |     |     |       |     |     |     | Leu | •   |
| M B        | , , , , , , | IIIC L | val  |       |     | voh | 116 | · •a: | 260 |     | , 101 | P   | _00 | 265 |     |     |
|            |             |        |      | 255   |     |     |     |       | 200 |     |       |     |     | 200 |     |     |

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| cca | gct | gga | aag | tac      | gtt | cgg  | agg  | gcg    | gag | aag | aag  | gcc | acc    | ctg  | tgt  | 869  |
|-----|-----|-----|-----|----------|-----|------|------|--------|-----|-----|------|-----|--------|------|------|------|
| Pro | Ala | Gly | Lys | Tyr      | Val | Arg  | Arg  | Ala    | Glu | Lys | Lys  | Ala | Thr    | Leu  | Cys  |      |
|     |     |     | 270 |          |     |      |      | 275    |     |     |      | •   | 280    |      |      | •    |
|     |     |     |     |          |     |      |      |        |     |     |      |     |        |      |      |      |
| cag | ctg | gag | gtg | gac      | gtg | ctg  | tgg  | tca    | gat | gtg | atc  | agt | cac    | çca  | ccg  | 917  |
| Gln | Leu | Glu | Val | Asp      | Val | Leu  | Trp  | Ser    | Asp | Val | ile  | Ser | His    | Pro  | Pro  |      |
|     |     | 285 |     |          |     |      | 290  |        |     |     |      | 295 |        |      |      |      |
|     |     |     |     |          |     |      |      |        |     |     |      |     |        |      |      |      |
| gag | ggg | cag | tgg | cag      | cgc | att  | gca  | CCC    | ctg | cgt | gtg  | ctt | agc    | ttc  | gac  | 965  |
| Glu | Gly | Gin | Trp | Gln      | Arg | He   | Ala  | Pro    | Leu | Arg | Val  | Leu | Ser    | Phe  | Asp  |      |
|     | 300 |     |     |          |     | 305  |      |        |     |     | 310  |     |        |      |      |      |
|     |     |     |     |          |     |      |      |        |     |     |      |     |        |      |      |      |
| atc | gag | tgt | gct | ggc      | cga | aaa  | ggc  | atc    | ttc | cct | gag  | cct | gag    | cgt  | gac  | 1013 |
| He  | Glu | Cys | Ala | Gly      | Arg | Lys  | Gly  | He     | Phe | Pro | Glu  | Pro | Glu    | Arg  | Asp  |      |
| 315 |     |     |     |          | 320 |      |      |        |     | 325 |      |     |        |      | 330  |      |
|     |     |     |     |          |     |      |      |        |     |     |      |     |        |      |      |      |
| CCC | gtg | atc | cag | atc      | tgt | tct  | ctg  | ggg    | ctg | cgc | tgg  | ggg | gag    | ccg  | gag  | 1061 |
|     |     |     | Gin |          |     |      |      |        |     |     |      |     |        |      |      |      |
|     |     |     |     | 335      |     |      |      | -      | 340 |     | •    |     |        | 345  |      |      |
|     |     |     |     |          |     |      |      |        |     |     |      |     |        |      |      |      |
| cca | ttc | ttg | cgt | ctg      | gca | ctc  | acg  | ctg    | cgg | CCC | tgt  | gcc | CCC    | atc  | ctg  | 1109 |
|     |     |     | Arg |          |     |      |      |        |     |     | _    |     |        |      | _    |      |
|     |     |     | 350 |          |     |      |      | 355    |     |     |      |     | 360    |      |      |      |
|     |     |     | 000 |          |     |      |      |        |     |     |      |     |        |      |      |      |
| øøt | gcc | aaa | gtg | CSE      | agc | tat  | gag  | cgg    | gaa | gaa | gac. | ctg | ctc    | Cag  | gcc  | 1157 |
|     | _   |     | Val |          |     |      |      |        |     |     |      |     |        |      |      | ,    |
| uıy | Λια | 365 |     | <b>u</b> | 001 |      | 370  |        | ۵.۵ | 4.4 | 7100 | 375 |        | ٠    |      |      |
|     |     |     |     |          |     |      | 0,0  |        |     |     |      | 0,0 |        |      |      |      |
| taa | acc | gac | ttc | ato      | ctt | gCC. | atø  | gac    | cct | gac | øtø  | atc | 800    | PPC. | tac  | 1205 |
|     |     |     | Phe |          |     |      |      |        |     |     |      |     |        |      |      | 1200 |
| ιτρ |     | vsh | LHE | 116      | LGu | 385  | MGE  | nop    | 110 | Agh | 390  | 110 | 114    | uly  | ı yı |      |
|     | 380 |     |     |          |     | 300  |      |        |     |     | 090  |     |        |      |      |      |
| 000 | a++ | ~~~ | aac | +++      | ave | 0+0  | cce  | tac    | oto | ato | tet  | Caa | ave    | Cac  | acc  | 1253 |
|     |     |     |     |          |     |      |      |        |     |     |      |     |        |      |      | 1200 |
|     | 118 | uin | Asn | FIRE     |     |      | Li O | ıyr    | Leu |     |      | M B | AId    | um   |      |      |
| 395 |     |     |     |          | 400 |      |      |        |     | 405 |      |     |        |      | 410  |      |
|     |     | •   |     |          | 4   |      |      | ۔ عارہ |     |     |      |     | -لىدىد | مطبع |      | 1001 |
| cta | aag | gtg | gac | cgc      | TTC | CCT  | TTC  | ctg    | ggc | cgc | gtg  | act | ggt    | CTC  | cgc  | 1301 |

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| Leu | Lys | Val | Asp        | Arg<br>415 | Phe | Pro | Phe | Leu | Gly<br>420 | Arg | Val | Thr | Gly | Leu<br>425 | Arg |      |
|-----|-----|-----|------------|------------|-----|-----|-----|-----|------------|-----|-----|-----|-----|------------|-----|------|
| tcc | aac | atc | cgt        | gac        | tcc | tcc | ttc | caa | tca        | agg | cag | gtc | ggc | cgg        | cgg | 1349 |
| Ser | Asn | He  | Arg        | Asp        | Ser | Ser | Phe | Gln | Ser        | Arg | Gln | Val | Gly | Arg        | Arg |      |
|     |     |     | 430        |            |     |     |     | 435 |            |     |     |     | 440 |            |     |      |
| gac | agt | aag | gtg        | atc        | agc | atg | gtg | ggt | cgc        | gtt | cag | atg | gat | atg        | ctg | 1397 |
| Asp | Ser | Lys | Val        | He         | Ser | Met | Val | Gly | Arg        | Val | Gln | Met | Asp | Met        | Leu |      |
|     |     | 445 |            |            |     |     | 450 |     |            |     |     | 455 |     |            |     |      |
| cag | gtg | ctg | ctt        | cgg        | gaa | cac | aag | ctc | cgc        | tcc | tac | acg | ctc | aac        | gct | 1445 |
| Gln | Val | Leu | Leu        | Arg        | Glu | His | Lys | Leu | Arg        | Ser | Tyr | Thr | Leu | Asn        | Ala |      |
|     | 460 |     |            |            |     | 465 |     |     |            |     | 470 |     |     |            |     |      |
|     |     |     |            |            |     | ggc |     |     |            |     |     |     |     |            |     | 1493 |
| Val | Ser | Phe | His        | Phe        |     | Gly | Glu | Gin | Lys        |     | Asp | Val | Gln | His        |     |      |
| 475 | •   |     |            |            | 480 |     |     |     |            | 485 |     |     |     |            | 490 |      |
|     |     |     |            |            |     |     |     |     |            |     |     |     |     |            | ctg | 1541 |
| He  | He  | Thr | Asp        |            | GIn | Asn | Gly | Asn |            | Gin | Inr | Arg | Arg |            |     |      |
|     |     |     |            | 495        |     |     |     |     | 500        |     |     |     |     | 505        |     | 1500 |
|     |     |     |            |            |     |     |     |     |            |     |     |     |     |            | gag | 1589 |
| Ala | Val | Tyr | Cys<br>510 |            | Lys | Asp | Ala | 515 |            | Pro | Leu | Arg | 520 |            | GIU |      |
| cgc | ctt | atg | gtg        | ctg        | gtg | aat | aat | gtg | gag        | atg | gcg | cgt | gto | acg        | ggt | 1637 |
|     |     |     |            |            |     |     |     |     |            |     |     |     |     |            | Gly |      |
|     |     | 525 |            |            |     |     | 530 |     |            |     |     | 535 |     |            |     |      |
| gta | CCC | ctt | ggg        | tac        | ctg | ctc | acc | cgg | ggc        | cag | cag | gto | aag | gto        | gtg | 1685 |
| Val | Pro | Leu | Gly        | Tyr        | Leu | Leu | Thr | Arg | Gly        | Gln | Gin | Val | Lys | Val        | Val |      |
|     | 540 |     |            |            |     | 545 |     |     |            |     | 550 |     |     |            |     |      |
| tct | cag | ctg | ctg        | cgc        | cag | gcc | atg | cgo | cag        | ggg | ctg | ctg | ate | cct        | gtg | 1733 |
| Ser | GIn | Leu | Leu        | Arg        | Gln | Ala | Met | Arg | GIn        | Gly | Leu | Leu | Met | . Pro      | Val |      |

| 555           | 560              | 565   | 570         |
|---------------|------------------|---|-------------|
|               |                  | ac tac acg gga gcc aca<br>sp Tyr Thr Gly Ala Thr<br>580       |             |
|               | y Tyr Tyr Asp Va | te ecc att gee acc etg<br>al Pro IIe Ala Thr Leu<br>595       |             |
|               | o Ser lie Met Mo | tg gcc cat aat ctg tgc<br>et Ala His Asn Leu Cys<br>10 615    |             |
|               |                  | ag aag ctg ggc ctt aaa<br>In Lys Leu Giy Leu Lys<br>630       |             |
|               |                  | at gag ttt gtg aag tca<br>sp Glu Phe Val Lys Ser<br>645       |             |
|               |                  | etg gag aat ctg ctg agt<br>eu Glu Asn Leu Leu Ser<br>660      |             |
| Arg Ala Lys A |                  | eag gag acg gac ccc ctg<br>Bin Glu Thr Asp Pro Leu<br>675     |             |
| -             | ly Arg Gin Leu A | gca cta aaa gtg agt gcc<br>Ala Leu Lys Val Ser Ala<br>690 695 | Asn Ser Val |
|               |                  | gtg ggc aag ctg cca tgt<br>Val Gly Lys Leu Pro Cys<br>710     |             |

|      | cag        | _                        | -                               |                          |                   |            |                          |                                 |                          |                   |            |                          |                          |                                 |                   | 2213 |
|------|------------|--------------------------|---------------------------------|--------------------------|-------------------|------------|--------------------------|---------------------------------|--------------------------|-------------------|------------|--------------------------|--------------------------|---------------------------------|-------------------|------|
|      | Gln        | Ser                      | Val                             | Thr                      |                   | Phe        | Gly                      | Arg                             | GIN                      |                   | He.        | Glu                      | Lys                      | ınr                             |                   |      |
| 715  |            |                          |                                 |                          | 720               |            |                          |                                 |                          | 725               |            |                          |                          |                                 | 730               |      |
| car  | ctt        | oto                      | gag                             | tee                      | aag               | .tac       | acc                      | ete                             | gaa                      | aat               | ggC        | tac                      | gat                      | acc                             | aac               | 2261 |
| -    | Leu        | _                        |                                 |                          |                   |            |                          |                                 |                          |                   |            |                          |                          |                                 |                   |      |
| uiii | Leu        | Vai                      | ulu                             | 735                      | Lyo               | 131        | 1111                     | 141                             | 740                      | AGII              | uıy        |                          | 700                      | 745                             | 71011             |      |
|      |            |                          |                                 | 700                      |                   |            |                          |                                 | , 10                     |                   |            |                          |                          |                                 |                   |      |
| gcc  | aag        | gta                      | gtc                             | tac                      | ggt               | gac        | acg                      | gac                             | tct                      | gtg               | atg        | tgc                      | cgg                      | ttt                             | ggc               | 2309 |
| Ala  | Lys        | Val                      | Val                             | Tyr                      | Gly               | Asp        | Thr                      | Asp                             | Ser                      | Val               | Met        | Cys                      | Arg                      | Phe                             | Gly               |      |
|      |            |                          | 750                             |                          |                   |            |                          | 755                             |                          |                   |            |                          | 760                      |                                 |                   |      |
|      |            |                          |                                 |                          |                   |            |                          |                                 |                          |                   |            |                          |                          |                                 |                   |      |
| gtc  | tcc        | tct                      | gtg                             | gct                      | gaa               | gca        | atg                      | tct                             | ctg                      | ggg               | cgg        | gag                      | gct                      | gca                             | aac               | 2357 |
| Val  | Ser        | Ser                      | Val                             | Ala                      | Glu               | Ala        | Met                      | Ser                             | Leu                      | Gly               | Arg        | Glu                      | Ala                      | Ala                             | Asn               |      |
|      |            | 765                      |                                 |                          |                   |            | 770                      |                                 |                          |                   |            | 775                      |                          |                                 |                   | •    |
|      | •          |                          |                                 |                          |                   |            |                          |                                 |                          |                   |            |                          |                          |                                 |                   |      |
| tgg  | gta        | tcc                      | agt                             | cac                      | ttc               | cca        | tca                      | CCC                             | atc                      | cgg               | ctg        | gag                      | ttc                      | gag                             | aag               | 2405 |
| Trp  | Val        | Ser                      | Ser                             | His                      | Phe               | Pro        | Ser                      | Pro                             | He                       | Arg               | Leu        | Glu                      | Phe                      | Glu                             | Lys               |      |
|      | 780        |                          |                                 |                          |                   | 785        |                          |                                 |                          |                   | 790        |                          |                          |                                 |                   |      |
|      |            |                          |                                 |                          |                   |            |                          |                                 |                          |                   |            |                          |                          |                                 |                   | •    |
| gtt  | tac        | ttc                      | cca                             | tac                      | ctg               | ctc        | atc                      | agc                             | aag                      | aag               | cgc        | tat                      | gct                      | ggc                             | ctg               | 2453 |
| Val  | Tyr        | Phe                      | Pro                             | Tyr                      | Leu               | Leu        | He                       | Ser                             | Lys                      | Lys               | Arg        | Tyr                      | Ala                      | Gly                             | Leu               |      |
| 795  |            |                          |                                 |                          | 800               |            |                          | •                               |                          | 805               |            |                          |                          |                                 | 810               |      |
|      |            |                          |                                 |                          |                   |            |                          |                                 |                          |                   |            |                          |                          |                                 |                   |      |
| ctc  | ttc        | tcc                      | tcc                             | cgc                      | tet               | orat.      | acc                      |                                 |                          |                   |            |                          |                          |                                 |                   | AFA4 |
| Leu  |            |                          |                                 | _                        |                   | gal        | goo                      | Gal                             | gac                      | aaa               | atg        | gac                      | tgc                      | aag                             | ggc               | 2501 |
|      | Phe        | Ser                      | Ser                             |                          |                   |            |                          |                                 | _                        | _                 |            |                          | _                        | _                               |                   | 2501 |
|      | Phe        | Ser                      | Ser                             |                          | Ser               |            |                          |                                 | _                        | _                 |            |                          | _                        | _                               |                   | 2501 |
|      | Phe        | Ser                      | Ser                             | Arg                      | Ser               |            |                          |                                 | Asp                      | _                 |            |                          | _                        | Lys                             |                   | 2501 |
| ctg  | Phe<br>gag |                          |                                 | Arg<br>815               | Ser               | Asp        | Ala                      | His                             | Asp<br>820               | Lys               | Met        | Asp                      | Cys                      | Lys<br>825                      | Gly               | 2549 |
|      |            | gct                      | gtg                             | Arg<br>815<br>cgc        | Ser               | Asp        | Ala                      | His<br>tgt                      | Asp<br>820<br>ccc        | Lys<br>ctg        | Met        | Asp                      | Cys                      | Lys<br>825<br>ctc               | Gly<br>gtt        |      |
|      | gag        | gct                      | gtg                             | Arg<br>815<br>cgc<br>Arg | Ser               | Asp        | Ala                      | His<br>tgt                      | Asp<br>820<br>ccc<br>Pro | Lys<br>ctg        | Met        | Asp                      | Cys                      | Lys<br>825<br>ctc               | Gly<br>gtt        |      |
| Leu  | gag<br>Glu | gct<br>Ala               | gtg<br>Val<br>830               | Arg<br>815<br>cgc<br>Arg | Ser<br>agg<br>Arg | gac<br>Asp | Ala<br>aac<br>Asn        | tgt<br>Cys<br>835               | Asp<br>820<br>ccc<br>Pro | Ctg<br>Leu        | gtg<br>Val | gcc<br>Ala               | aac<br>Asn<br>840        | Lys<br>825<br>ctc<br>Leu        | Gly<br>gtt<br>Val | 2549 |
| Leu  | gag<br>Glu | gct<br>Ala               | gtg<br>Val<br>830<br>ctg        | Arg<br>815<br>cgc<br>Arg | agg<br>Arg        | gac<br>Asp | Ala<br>aac<br>Asn        | tgt<br>Cys<br>835               | Asp<br>820<br>ccc<br>Pro | ctg<br>Leu<br>cgg | gtg<br>Val | gcc<br>Ala               | aac<br>Asn<br>840<br>gat | Lys<br>825<br>ctc<br>Leu<br>ggg | gtt<br>Val        |      |
| Leu  | gag<br>Glu | gct<br>Ala<br>tct<br>Ser | gtg<br>Val<br>830<br>ctg<br>Leu | Arg<br>815<br>cgc<br>Arg | agg<br>Arg        | gac<br>Asp | aac<br>Asn<br>ctc<br>Leu | tgt<br>Cys<br>835<br>gtg<br>Val | Asp<br>820<br>ccc<br>Pro | ctg<br>Leu<br>cgg | gtg<br>Val | gcc<br>Ala<br>cct<br>Pro | aac<br>Asn<br>840<br>gat | Lys<br>825<br>ctc<br>Leu<br>ggg | gtt<br>Val        | 2549 |
| Leu  | gag<br>Glu | gct<br>Ala               | gtg<br>Val<br>830<br>ctg<br>Leu | Arg<br>815<br>cgc<br>Arg | agg<br>Arg        | gac<br>Asp | Ala<br>aac<br>Asn        | tgt<br>Cys<br>835<br>gtg<br>Val | Asp<br>820<br>ccc<br>Pro | ctg<br>Leu<br>cgg | gtg<br>Val | gcc<br>Ala               | aac<br>Asn<br>840<br>gat | Lys<br>825<br>ctc<br>Leu<br>ggg | gtt<br>Val        | 2549 |

|             |                   | to tog gad otg otg                         |                    | 645       |
|-------------|-------------------|--|--------------------|-----------|
|             |                   | le Ser Asp Leu Leu (                       | Cys Asn Arg Ile    |           |
| 860         | 865               | 870 <sub>.</sub>                           |                    |           |
| gac atc tcc | cag ctg gtc atc a | cc aaa gag ttg acc                         | cgc gca gca gca 20 | 693       |
| _           |                   | nr Lys Glu Leu Thr                         |                    |           |
| 875         | 880               | 885  | 890                |           |
|             |                   | ·  |                    |           |
| _           |                   | ac gtg gag ctg gct                         | 0-0 -000           | 741       |
| Asp Tyr Ala |                   | is Val Glu Leu Ala                         |                    |           |
|             | 895               | 900  | 905                |           |
| aag cgc gac | ccc ggc agt gcg c | cc agc ctg ggt gac                         | cga gtc ccc tat 2  | 789       |
| _           |                   | ro Ser Leu Gly Asp                         |                    |           |
|             | 910               | 915  | 920                |           |
|             |                   |  |                    |           |
|             |                   | gt gtg gcc gcc tac                         |                    | 2837      |
|             |                   | ily Val Ala Ala Tyr<br>30                  | 935                |           |
| 925         | 3                 | ,  | 333                |           |
| gac ccc ctg | ttt gtg ctg gag c | ac ago otg coc atc                         | gac act cag tac 2  | 2885      |
| Asp Pro Leu | Phe Val Leu Glu H | lis Ser Leu Pro Ile                        | Asp Thr Gin Tyr    |           |
| 940         | 945               | 950  |                    |           |
|             |                   |  |                    | 1022      |
|             |                   | ag ccg ctc ttg cgc<br>ys Pro Leu Leu Arg   |                    | 2933      |
| 955         | 960               | 965  | 970                |           |
| 560         | 000               |  |                    |           |
| atc ctg ggt | gag ggc cgt gca g | gag tot gtg ctg ctg                        | cgc ggt gac cac 2  | 2981      |
| lle Leu Gly | Glu Gly Arg Ala G | ilu Ser Val Leu Leu                        |                    |           |
|             | 975               | 980  | 985                |           |
|             |                   | ore ore sta are                            | age etc. ttg gee.  | 3029      |
|             |                   | acc agc aag gtg ggc<br>Thr Ser Lys Val Gly |                    | . <u></u> |
| iii nig Oya | 990               | 995  | 1000               |           |
|             |                   |  |                    |           |
| ttc acc aag | cgc cgc aac tgt   | tgc att ggc tgc cg                         | go too gta ato     | 3074      |

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| Phe  | Thr    | Lys<br>1005 | Arg  | Arg | Asn | Cys  | Cys<br>1010 | He         | Gly | Cys                                     | Arg        | Ser<br>1015 | Val       | lle  |             |
|------|--------|-------------|------|-----|-----|------|-------------|------------|-----|---|------------|-------------|-----------|------|-------------|
| gac  | cat    | caa         | gga  | gcc | gtg | tgt  | aag         | ttc        | tgt | cag                                     | cca        | cgg         | gag       | tcg  | 3119        |
| Asp  | His    | Gln         | Gly  | Ala | Val | Cys  | Lys         | Phe        | Cys | Gln                                     | Pro        | Arg         | Glu       | Ser  |             |
|      |        | 1020        |      |     |     |      | 1025        |            |     |   |            | 1030        |           |      |             |
| gog. | oto    | <b>t</b> ot | 000  | 224 | nen | ata  | tca         | 020        | cta | 22+                                     | «cc        | <b>+</b> +a | поо       | gaa. | 3164        |
|      |        |             | -    |     |     |      | Ser         |            |     |   |            |             |           |      | 9104        |
| uiu  | LCu    | 1035        | um   | LyG | uiu | ·u,  | 1040        | 1110       | LUG | ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,, | ,,,,,      | 1045        | aiu       | u.u  |             |
|      |        | 1000        |      |     |     |      | 10.0        |            |     |   |            |             |           |      |             |
| cgg  | ttc    | tct         | cgc  | ctc | tgg | aca  | cag         | tgt        | caa | cgc                                     | tgc        | cag         | ggc       | agc  | 3209        |
| Arg  | Phe    | Ser         | Arg  | Leu | Trp | Thr  | Gln         | Cys        | Gin | Arg                                     | Cys        | Gln         | Gly       | Ser  |             |
|      |        | 1050        |      |     |     |      | 1055        |            |     |   |            | 1060        |           |      |             |
|      |        |             |      |     |     |      |             |            |     |   |            |             |           |      |             |
| ttg  | cat    | gag         | gac  | gtc | atc | tgt  | acc         | agc        | cgt | gac                                     | tgt        | CCC         | atc       | ttc  | 3254        |
| Leu  | His    | Glu         | Asp  | Val | He  | Cys  | Thr         | Ser        | Arg | Asp                                     | Cys        |             | He        | Phe  |             |
|      |        | 1065        |      |     |     |      | 1070        |            |     |   |            | 1075        |           |      |             |
| +    | a-b-a- |             |      |     | ~+~ | 000  |             | <b>~~~</b> | ota | ~~~                                     | <b>~~~</b> | 007         | <b>~~</b> | 000  | 3299        |
|      | _      | _           | _    | _   |     | _    | aag<br>Lys  | -          | _   | _                                       | -          | _           | _         |      | <b>3233</b> |
| ıyı  | Mec    | 1080        | Lys  | Lys | Vai | AI g | 1085        | nsp        | Leu | uiu                                     | voh        | 1090        | uiu       | AI S |             |
|      |        | 1000        |      |     |     |      | 1000        |            |     |   |            | 1030        |           |      |             |
| ctg  | ctg    | cag         | cgc  | ttt | gga | ccg  | CCC         | ggc        | cct | gag                                     | gcc        | tgg         | tga       |      | 3341        |
| Leu  | Leu    | Gln         | Arg  | Phe | Gly | Pro  | Pro         | Gly        | Pro | Glu                                     | Ala        | Trp         |           |      |             |
|      |        | 1095        |      |     |     |      | 1100        |            |     |   |            | 1105        |           |      |             |
| ŧ    |        |             |      |     |     |      |             |            |     |   |            |             |           |      |             |
| cct  | gaca   | ogg ga      | acaa | g   |     |      |             |            |     |   |            |             |           |      | 3357        |
|      |        |             |      |     |     |      |             |            |     |   |            |             |           |      |             |

`**<210> 87** 

<211> 1105

<212> PRT

<213> Mus musculus

<400> 87

Met Asp Cys Lys Arg Arg Gin Gly Pro Gly Pro Giy Val Pro Pro Lys 5 Arg Ala Arg Gly His Leu Trp Asp Glu Asp Glu Pro Ser Pro Ser Gln 20 25 30 Phe Glu Ala Asn Leu Ala Leu Leu Glu Glu Ile Glu Ala Glu Asn Arg Leu Gin Giu Ala Giu Giu Leu Gin Leu Pro Pro Giu Giy Thr Val 50 55 60 Gly Gly Gln Phe Ser Thr Ala Asp Ile Asp Pro Arg Trp Arg Arg Pro 80 65 · 70 Thr Leu Arg Ala Leu Asp Pro Ser Thr Glu Pro Leu ile Phe Gin Gin 85 95 Leu Glu IIe Asp His Tyr Val Gly Ser Ala Pro Pro Leu Pro Glu Gly 110 100 105 Pro Leu Pro Ser Arg Asn Ser Val Pro IIe Leu Arg Ala Phe Gly Val 125 120 115 Thr Asp Glu Gly Phe Ser Val Cys Cys His Ile Gln Gly Phe Ala Pro 130 135 140

Tyr Phe Tyr Thr Pro Ala Pro Pro Gly Phe Gly Ala Glu His Leu Ser

| 145            |            |            |            | 150        |            |            |            |            | 155        |            |            |            |            | 160        |
|----------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| Glu Leu        | Gln        | GIn        | Glu<br>165 | Leu        | Asn        | Ala        | Ala        | lle<br>170 | Ser        | Arg        | Asp        | Gln        | Arg<br>175 | Gly        |
| Gly Lys        | Glu        | Leu<br>180 | Ser        | Gly        | Pro        | Ala        | Vai<br>185 | Leu        | Ala        | He         | Glu        | Leu<br>190 |            | Ser        |
| Arg Glu        | Ser<br>195 | Met        | Phe        | Gly        | Tyr        | His<br>200 | Gly        | His        | Gly        | Pro        | Ser<br>205 | Pro        | Phe        | Leu        |
| Arg lle<br>210 | Thr        | Leu        | Ala        | Leu        | Pro<br>215 | Arg        | Leu        | Met        | Ala        | Pro<br>220 | Ala        | Arg        | Arg        | Leu        |
| Leu Glu<br>225 | Gin        | Gly        | Val        | Arg<br>230 | Val        | Pro        | Gly        | Leu        | Gly<br>235 | Thr        | Pro        | Ser        | Phe        | Ala<br>240 |
| Pro Tyr        | Glu        | .Ala       | Asn<br>245 | Val        | Asp        | Phe        | Glu        | ile<br>250 | Arg        | Phe        | Met        | Val        | Asp<br>255 | Ala        |
| Asp lie        | Val        | Gly<br>260 |            | Asn        | Trp        | Leu        |            |            |            | Ala        |            | Lys<br>270 | Tyr        | Val        |
| Arg Arg        | Ala<br>275 |            | Lys        | Lys        | Ala        | Thr<br>280 | Leu        | Cys        | Gln        | Leu        | Glu<br>285 | Val        | Asp        | Val        |
| Leu Trp<br>290 |            | Asp        | Val        | He         | Ser<br>295 | His        | Pro        | Pro        | Glu        | Gly<br>300 | Gln        | Trp        | Gln        | Arg        |

| lle<br>305 | Ala        | Pro        | Leu        | Arg         | Va I<br>310 | Leu        | Ser        | Phe        | Asp        | l le<br>315 | Glu <sub>.</sub> | Cys        | Ala        | Gly        | Arg<br>320 |
|------------|------------|------------|------------|-------------|-------------|------------|------------|------------|------------|-------------|------------------|------------|------------|------------|------------|
| Lys        | Gly        | lle        | Phe        | Pro<br>325  | Glu         | Pro        | Glu        | Arg        | Asp<br>330 | Pro         | Val              | lle        | GIn        | 11e<br>335 | Cys        |
| Ser        | Leu        | Gly        | Leu<br>340 | Arg         | Trp         | Gly        | Glu        | Pro<br>345 |            | Pro         | Phe              | Leu        | Arg<br>350 | Leu        | Ala        |
| Leu        | Thr        | Leu<br>355 |            | Pro         | Cys         | Ala        | Pro<br>360 | lle        | Leu        | Gly         | Ala              | Lys<br>365 | Val        | Gln        | Ser        |
| Tyr        | Glu<br>370 |            | Glu        | Glu         | Asp         | Leu<br>375 |            | GIn        | Ala        | Trp         | Ala<br>380       | Asp        | Phe        | lle        | Leu        |
| Ala<br>385 |            | Asp        | Pro        | Asp         | Va I<br>390 |            | Thr        | Gly        | Tyr        | Asn<br>395  |                  | Gln        | Asn        | Phe        | Asp<br>400 |
| Leu        | Pro        | Tyr        | Leu        | l le<br>405 |             | Arg        | Ala        | GIn        | Ala<br>410 |             | Lys              | Val        | Asp        | Arg<br>415 |            |
| Pro        | Phe        | Leu        | Gly<br>420 |             | ; Val       | Thr        | Gly        | Leu<br>425 |            | Ser         | Asn              | He         | Arg<br>430 |            | Ser        |
| Ser        | · Phe      | . Gln      | . Ser      | · Arg       | Gln         | Val        | Glv        | Are        | : Arg      | Asp         | Ser              | Lys        | : Val      | lle        | Ser        |

440

435

445

| Met | Val | Gly | Arg | Val | Gin | Met | Asp | Met | Leu | Gln | Val | Leu | Leu | Arg | Glu |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
|     | 450 |     |     |     |     | 455 |     |     |     |     | 460 |     |     |     |     |

His Lys Leu Arg Ser Tyr Thr Leu Asn Ala Val Ser Phe His Phe Leu 465 470 475 480

Gly Glu Gln Lys Glu Asp Val Gln His Ser IIe IIe Thr Asp Leu Gln 485 490 495

Asn Gly Asn Glu Gln Thr Arg Arg Leu Ala Val Tyr Cys Leu Lys 500 505 510

Asp Ala Phe Leu Pro Leu Arg Leu Leu Glu Arg Leu Met Val Leu Val 515 520 525

Asn Asn Val Glu Met Ala Arg Val Thr Gly Val Pro Leu Gly Tyr Leu 530 535 540

Leu Thr Arg Gly Gln Gln Val Lys Val Val Ser Gln Leu Leu Arg Gln 545 550 555 560

Ala Met Arg Gin Gly Leu Leu Met Pro Val Val Lys Thr Glu Gly Ser 565 570 575

Glu Asp Tyr Thr Gly Ala Thr Val lle Glu Pro Leu Lys Gly Tyr Tyr 580 585 590

| Asp        | Val        | Pro<br>595 | He         | Ala        | Thr        | Leu        | Asp<br>600 | Phe        | Ser        | Ser               |            | Tyr<br>605 | Pro<br>·   | Ser        | lle        |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------------|------------|------------|------------|------------|------------|
| Met        | Met<br>610 | Ala        | His        | Asn        | Leu        | Cys<br>615 | Tyr        | Thr        | Thr        | Leu               | Leu<br>620 | Arg        | Pro        | Gly        | Ala        |
| Ala<br>625 | GIn        | Lys        | Leu        | Gly        | Leu<br>630 | Lys        | Pro        | Asp        | Glu        | Phe<br><b>635</b> | He         | Lys        | Thr        | Pro        | Thr<br>640 |
| Gly        | Asp        | Glu        | Phe        | Val<br>645 | Lys        | Ser        | Ser        | Val        | Arg<br>650 | Lys               | Gly        | Leu        | Leu        | Pro<br>655 | GIn        |
| He         | Leu        | Glu        | Asn<br>660 | Leu        | Leu        | Ser        | Ala        | Arg<br>665 | Lys        | Arg               | Ala        | Lys        | Ala<br>670 | Glu        | Leu        |
| Ala        | Gln        | Glu<br>675 | Thr        | Asp        | Pro        | Leu        | Arg<br>680 | Arg        | GIn        | Val               | Leu        | Asp<br>685 | Gly        | Arg        | Gln        |
| Leu        | Ala<br>690 |            | Lys        | Val        | Ser        | Ala<br>695 | Asn        | Ser        | Val        | Tyr               | Gly<br>700 | Phe        | Thr        | Gly        | Ala        |
| GIn<br>705 |            | Gly        | Lys        | Leu        | Pro<br>710 | Cys        | Leu        | Glu        | lle        | Ser<br>715        | -          | Ser        | Val        | Thr        | Gly<br>720 |
| Phe        | Gly        | Arg        | Gln        | Met<br>725 |            | Glu        | Lys        | Thr        | Lys<br>730 | Gln               | Leu        | Va!        | Glu        | Ser<br>735 | Lys        |

Tyr Thr Val Glu Asn Gly Tyr Asp Ala Asn Ala Lys Val Val Tyr Gly
740 745 750

Asp Thr Asp Ser Val Met Cys Arg Phe Gly Val Ser Ser Val Ala Glu
755 760 765

Ala Met Ser Leu Gly Arg Glu Ala Ala Asn Trp Val Ser Ser His Phe 770 775 780

Pro Ser Pro 11e Arg Leu Glu Phe Glu Lys Val Tyr Phe Pro Tyr Leu 785 790 795 800

Leu lie Ser Lys Lys Arg Tyr Ala Gly Leu Leu Phe Ser Ser Arg Ser 805 810 815

Asp Ala His Asp Lys Met Asp Cys Lys Gly Leu Glu Ala Val Arg Arg 820 825 830

Asp Asn Cys Pro Leu Val Ala Asn Leu Val Thr Ser Ser Leu Arg Arg 835 840 845

lle Leu Val Asp Arg Asp Pro Asp Gly Ala Val Ala His Ala Lys Asp 850 855 860

Val lle Ser Asp Leu Leu Cys Asn Arg lle Asp lle Ser Gln Leu Val 865 870 875 880

lle Thr Lys Glu Leu Thr Arg Ala Ala Ala Asp Tyr Ala Gly Lys Gln

|               |                  | 885                  | 890                  | )                     | 895                 |
|---------------|------------------|----------------------|----------------------|-----------------------|---------------------|
| Ala Hi        | s Val Glu<br>900 |                      | Arg Met Arg<br>905   | Lys Arg Asp           | Pro Gly Ser<br>910  |
| Ala Pr        | o Ser Leu<br>915 | ı Gly Asp Arg        | ; Val Pro Tyr<br>920 | Val IIe IIe<br>925    | Gly Ala Ala         |
| Lys GI<br>93  |                  | ı Ala Tyr Met<br>935 |                      | Asp Pro Leu<br>940    | Phe Val Leu         |
| Glu Hi<br>945 | s Ser Leu        | ıPro lle Asp<br>950  | Thr Gln Tyr          | Tyr Leu Glu<br>955    | Gin Gin Leu<br>960  |
| Ala Ly        | s Pro Leu        | ıLeu Arg ile<br>965  | Phe Glu Pro<br>970   | ile Leu Gly           | Glu Gly Arg<br>975  |
| Ala Gi        | u Ser Val<br>980 |                      | Gly Asp His<br>985   | Thr Arg Cys           | Lys Thr Val<br>990  |
| Leu Th        | r Ser Lys<br>995 | s Val Gly Gly        | Leu Leu Al           | a Phe Thr Lys         | s Arg Arg Asn<br>05 |
|               | s lle Gi<br>10   |                      | er Vallle A<br>915   | sp His Gln (<br>1020  | Gly Ala Val         |
| _             | s Phe Cy<br>25   |                      | g Glu Ser 6<br>30    | ilu Leu Tyr (<br>1035 | Gin Lys Giu         |

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Val Ser His Leu Asn Ala Leu Glu Glu Arg Phe Ser Arg Leu Trp 1040 1045 1050

Thr Gln Cys Gln Arg Cys Gln Gly Ser Leu His Glu Asp Val IIe 1055 1060 1065

Cys Thr Ser Arg Asp Cys Pro lle Phe Tyr Met Arg Lys Lys Vai 1070 1075 1080

Arg Lys Asp Leu Glu Asp Gin Glu Arg Leu Leu Gin Arg Phe Gly 1085 1090 1095

Pro Pro Gly Pro Glu Ala Trp 1100 1105

<210> 88

WO 2004/087960

<211> 3318

<212> DNA

<213> Mus musculus

<220>

<221> CDS

<222> (1).. (3318)

**<400> 88** 

atg gat tgt aag cgg cga caa gga cca ggc cct ggg gtg ccc cca aag Met Asp Cys Lys Arg Arg Gin Giy Pro Giy Pro Giy Vai Pro Pro Lys 48

1 5 10 15

| cgg  | gct | cga      | ggg  | cac | ctc | tgg | gat | gag  | gac | gag | cct | tcg | ccg  | tcg  | cag | 96  |
|------|-----|----------|------|-----|-----|-----|-----|------|-----|-----|-----|-----|------|------|-----|-----|
| Arg  | Ala | Arg      | Gly  | His | Leu | Trp | Asp | Glu  | Asp | Glu | Pro | Ser | Pro  | Ser  | Gln |     |
|      |     |          | 20   |     |     |     |     | 25   |     |     |     |     | 30   |      |     |     |
|      |     |          |      |     |     |     |     |      |     |     |     |     |      |      |     |     |
| ttt  | gag | gcg      | aac  | ctg | gca | ctg | ctg | gag  | gaa | ata | gag | gct | gag  | aac  | cgg | 144 |
|      |     |          |      |     |     | Leu |     |      |     |     |     |     |      |      |     | •   |
|      |     | 35       |      |     |     |     | 40  |      |     |     |     | 45  |      |      |     |     |
|      |     |          |      |     |     |     |     |      |     |     |     |     |      |      |     |     |
| ctg  | cag | gag      | gca  | gag | gag | gag | ctg | cag  | ctg | CCC | cca | gag | ggc  | acc  | gtg | 192 |
|      |     |          |      |     |     | Glu |     |      |     |     |     |     |      |      |     |     |
|      | 50  |          |      |     |     | ·55 |     |      |     |     | 60  |     |      |      |     |     |
|      |     |          |      |     |     |     |     |      |     |     | •   |     |      |      |     |     |
| ppt  | ggg | cag      | ttt  | tcc | act | gca | gac | att  | gac | cct | cgg | tgg | cgg  | cgg  | CCC | 240 |
|      |     |          |      |     |     | Ala |     |      |     |     |     |     |      |      |     |     |
| 65   | u., | <b>u</b> |      | 00. | 70  |     |     |      |     | 75  |     |     |      |      | 80  |     |
| 00   |     |          |      |     | 70  |     |     |      |     |     |     |     |      |      |     |     |
| 200  | ota | out      | acc. | cta | asc | CCC | 900 | aca  | asa | ccc | ctc | atc | ttc  | CSE  | Cag | 288 |
|      |     |          |      |     |     | Pro |     |      |     |     |     |     |      |      |     | 200 |
| 1111 | Leu | M B      | Mia  |     | voh | 710 | 961 | 1111 | 90  | 110 | LUU | 110 | 1110 | 95   | u   |     |
|      |     |          |      | 85  |     |     |     |      | 30  |     |     |     |      | 30   |     |     |
|      |     | بالباد   |      |     | 4.4 |     |     | too  | ~~~ | 000 | 000 | ota | 002  | maa. | aaa | 336 |
|      |     |          |      |     |     | gtg |     |      |     |     |     |     |      |      |     | 000 |
| Leu  | Glu | He       |      |     | ıyr | Val | шу  |      |     | Pro | Pro | Leu |      |      | шу  |     |
|      |     |          | 100  |     |     |     |     | 105  |     |     |     |     | 110  |      |     |     |
|      |     |          |      |     |     | _   |     |      |     |     |     |     |      |      |     | 004 |
|      |     |          |      |     |     | tca |     |      |     |     |     |     |      |      |     | 384 |
| Pro  | Leu |          |      | Arg | Asn | Ser |     |      | He  | Leu | Arg |     |      | Gly  | vai |     |
|      |     | 115      |      |     |     |     | 120 |      |     |     |     | 125 |      |      |     |     |
|      |     |          |      |     |     |     |     |      |     |     |     |     |      |      |     |     |
|      |     |          |      |     |     | gtc |     |      |     |     |     |     | •    |      |     | 432 |
| Thr  | Asp | Glu      | Gly  | Phe | Ser | Val | Cys | Cys  | His | He  | Gin | Gly | Phe  | Ala  | Pro |     |
|      | 130 |          |      |     |     | 135 |     |      |     |     | 140 | 1   |      |      |     |     |
|      |     |          |      |     |     |     |     |      |     |     |     |     |      |      |     |     |
| tac  | ttc | tac      | acc  | ccc | gcg | cct | cct | ggt  | ttt | ggg | gcc | gag | cac  | ctg  | agt | 480 |
| Tyr  | Phe | Tyr      | Thr  | Pro | Ala | Pro | Pro | Gly  | Phe | Gly | Ala | Glu | His  | Leu  | Ser |     |
| 145  |     |          |      |     | 150 | )   |     |      |     | 155 | i   |     |      |      | 160 |     |
|      |     |          |      |     |     |     |     |      |     |     |     |     |      |      |     |     |

|     |       |     |     |     | ctg<br>Leu |     |      |     |     |     |         |       |     |     |     | 528   |
|-----|-------|-----|-----|-----|------------|-----|------|-----|-----|-----|---------|-------|-----|-----|-----|-------|
|     |       |     |     | 165 |            |     |      |     | 170 | •   |         |       |     | 175 |     |       |
| ggg | aag   | gag | ctc | tca | ggg        | ccg | gca  | gtg | ctg | gca | ata     | gag   | cta | tgc | tcc | 576   |
| Gly | Lys   | Glu | Leu | Ser | Gly        | Pro | Ala  | Val | Leu | Ala | He      | Glu   | Leu | Cys | Ser |       |
|     |       |     | 180 |     |            |     |      | 185 |     |     |         |       | 190 |     |     |       |
| cgt | gag   | agc | atg | ttt | ggg        | tac | cac  | ggt | cat | ggc | cct     | tct   | cca | ttt | ctc | 624   |
| Arg | Glu   | Ser | Met | Phe | Gly        | Tyr | His  | Gly | His | Gly | Pro     | Ser   | Pro | Phe | Leu |       |
|     |       | 195 |     |     |            |     | 200  |     |     |     |         | 205   |     |     |     |       |
| cgc | atc   | acc | ctg | gca | cta        | ccc | cgc  | ctt | atg | gca | cca     | gcc   | cgc | cgc | ctt | 672   |
| Arg | He    | Thr | Leu | Ala | Leu        | Pro | Arg  | Leu | Met | Ala | Pro     | Ala   | Arg | Arg | Leu |       |
|     | 210   |     |     |     |            | 215 |      |     |     |     | 220     |       |     |     |     |       |
| ctg | gaa   | cag | ggt | gtc | cga        | gtg | cca  | ggc | ctg | ggc | acc     | ccg   | agc | ttc | gca | 720   |
| Leu | Glu   | Gin | Gly | Val | Arg        | Val | Pro  | Gly | Leu | Gly | Thr     | Pro   | Ser | Phe | Ala |       |
| 225 |       |     |     |     | 230        |     |      |     |     | 235 |         |       |     |     | 240 |       |
|     |       |     |     |     |            |     |      |     |     |     | ح باساد |       |     | ~~+ | ~~+ | 768   |
|     |       |     | •   |     | gtg        |     |      |     |     |     |         |       |     |     |     | 700   |
| Pro | ıyr   | uiu | Ala | 245 | Val        | wsh | Lile | uiu | 250 |     | THE     | ING L | Vai | 255 |     |       |
|     |       |     |     | 270 |            |     |      |     | 200 |     |         |       |     |     |     |       |
| gac | att   | gtg | gga | tgc | aac        | tgg | ttg  | gag | ctg | cca | gct     | gga   | aag | tac | gtt | 816   |
| Asp | He    | Val | Gly | Cys | Asn        | Trp | Leu  | Glu | Leu | Pro | Ala     | Gly   | Lys | Tyr | Val |       |
|     |       |     | 260 |     |            |     |      | 265 |     |     |         |       | 270 | ı   | •   |       |
|     |       |     |     |     |            |     |      |     |     |     |         |       |     |     |     | 004   |
| -60 | agg   | 0-0 |     |     |            |     |      |     |     |     |         |       |     |     | gtg | . 864 |
| Arg | Arg   |     |     | Lys | Lys        | Ala |      |     | Gys | GIN | Leu     |       |     | Asp | vai |       |
|     |       | 275 | 1   |     |            | -   | 280  | l   |     |     |         | 285   | ,   |     |     |       |
| cte | tgg   | tca | gat | gte | ato        | agt | cac  | cca | CCg | gag | ggg     | cag   | tgg | cag | cgc | 912   |
| Leu | ı Trp | Ser | Asp | Val | He         | Ser | His  | Pro | Pro | Glu | Gly     | Glr   | Trp | Gln | Arg |       |
|     | 290   | )   |     |     |            | 295 |      |     |     |     | 300     | )     |     |     |     |       |
|     |       |     |     |     |            |     |      |     |     |     |         |       |     |     |     |       |
| att | gca   | CCC | ctg | cgt | : gtg      | ctt | ago  | ttc | gac | ato | gag     | tgt   | gct | ggo | cga | 960   |

| l le<br>305 | Ala        | Pro        | Leu | Arg | Va I<br>310 | Leu        | Ser        | Phe | Asp | l le<br>315 | Glu  | Cys        | Ala | Gly | Arg<br>320 |      |
|-------------|------------|------------|-----|-----|-------------|------------|------------|-----|-----|-------------|------|------------|-----|-----|------------|------|
|             |            |            |     |     |             | cct<br>Pro |            |     |     |             |      |            |     |     | _          | 1008 |
|             |            |            |     | 325 |             |            |            |     | 330 |             |      |            |     | 335 |            |      |
|             |            |            |     |     |             | ggg        |            |     |     |             |      |            |     |     |            | 1056 |
| ser         | Leu        | шу         | 340 | Arg | пр          | Gly        | aiu        | 345 | aiu | rio         | FIIG | Leu        | 350 | LCu | AIG        |      |
|             |            |            |     |     |             | gcc        |            |     |     |             |      |            |     |     |            | 1104 |
| Leu         | Thr        | 355        | Arg | Pro | Cys         | Ala        | 360        | ile | Leu | Gly         | Ala  | 365        | vai | GIN | ser        |      |
|             | -          |            |     |     |             | ctg        |            |     |     |             |      |            |     |     |            | 1152 |
| Tyr         | Glu<br>370 | Arg        | Glu | Glu | Asp         | Leu<br>375 | Leu        | GIn | Ala | Irp         | 380  | Asp        | Phe | He  | Leu        |      |
|             |            |            |     |     |             | atc        |            |     |     |             |      |            |     |     |            | 1200 |
| Ala<br>385  | Met        | Asp        | Pro | Asp | 390         | lle        | Thr        | Gly | lyr | 4sn<br>395  | He   | Gin        | Asn | Pne | 400        |      |
|             |            |            |     |     |             | cgg        |            |     |     | _           | _    |            | _   | _   |            | 1248 |
| Leu         | Pro        | Tyr        | Leu | 405 | Ser         | Arg        | Ala        | Gin | 410 | Leu         | Lys  | Vai        | Asp | 415 | Pne        |      |
|             |            |            |     |     |             | act        |            |     |     |             |      |            |     |     |            | 1296 |
| Pro         | Phe        | Leu        | 420 | Arg | Val         | Thr        | GIY        | 425 | Arg | Ser         | Asn  | 11e        | 430 | Asp | ser        |      |
|             |            |            |     |     |             | gtc        |            |     |     |             |      |            |     |     |            | 1344 |
| Ser         | Phe        | GIn<br>435 |     | Arg | Gin         | Val        | Gly<br>440 | Arg | Arg | Asp         | Ser  | Lys<br>445 |     | lle | Ser        |      |
|             |            |            |     |     |             | atg        |            |     |     |             |      |            |     |     |            | 1392 |
| wet         | val        | uly        | Arg | val | uin         | Met        | ASP        | wet | Leu | นเท         | vai  | Leu        | Leu | arg | uıu        |      |

| 450                 | 455             | 46              | 60                  |      |
|---------------------|-----------------|-----------------|---------------------|------|
|                     |                 |                 |                     |      |
| cac aag ctc cgc tcc | c tac acg ctc a | ac gct gtg ag   | st tto cac tto ctg  | 1440 |
| His Lys Leu Arg Ser | r Tyr Thr Leu A | Asn Ala Vai Se  | er Phe His Phe Leu  |      |
| 465                 | 470             | 475             | 480                 |      |
|                     |                 |                 |                     |      |
| ggc gag cag aag gag | g gac gtt cag o | cac ago ato at  | to acc gac ctg cag  | 1488 |
| Gly Glu Gln Lys Glu |                 |                 |                     |      |
| 48                  |                 | 490             | 495                 |      |
|                     |                 | •               |                     |      |
| aat ggg aac gaa ca  | g acg cgc cgc ( | ogo otg goo gt  | tg tac tgc ctg aag  | 1536 |
| Asn Gly Asn Glu Gli |                 |                 |                     |      |
| 500                 |                 | 505             | 510                 |      |
|                     |                 |                 |                     |      |
| gac gcc ttt ctg cc  | a ctc cga cta   | cta gag cgc ci  | tt atg gtg ctg gtg  | 1584 |
| Asp Ala Phe Leu Pr  |                 |                 |                     |      |
| 515                 | 520             | _               | 525                 |      |
| ,010                |                 |                 |                     |      |
| aat aat oto oao at  | e ece cet etc   | ace eet eta c   | cc ctt ggg tac ctg  | 1632 |
| Asn Asn Val Glu Me  |                 |                 |                     |      |
| 530                 | 535             |                 | 40                  |      |
| 550                 | 000             | _               | ,                   |      |
| oto oco oga ago ca  | or can ote aan  | ate ata tet e   | ag ctg ctg cgc cag  | 1680 |
|                     |                 |                 | in Leu Leu Arg Gin  |      |
|                     | 550             | 555             | 560                 |      |
| 545                 | 550             | 000             |                     |      |
| ota oca ma          | ra eta eta eta  | eet ata ata a   | ag acc gag ggc agt  | 1728 |
| =                   |                 |                 | ys Thr Glu Gly Ser  | 5    |
|                     |                 | 570             | 575                 |      |
| 56                  | 00              | 370             | 070                 |      |
|                     |                 |                 |                     | 1776 |
|                     |                 |                 | etc aaa ggg tac tat | 1770 |
|                     | IY Ala Inr vai  |                 | eu Lys Gly Tyr Tyr  |      |
| 580                 |                 | 585             | 590                 |      |
|                     | _               | المناف بيوانيون |                     | 1004 |
|                     |                 |                 | ttg tac cca tcc atc | 1824 |
|                     |                 | rne Ser Ser L   | Leu Tyr Pro Ser Ile |      |
| 595                 | 600             |                 | 605                 |      |

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| _ |     |     |     |     |     | tgc<br>Cys<br>615 |  |     |     |     |  |     |     | 1872 |
|---|-----|-----|-----|-----|-----|-------------------|--|-----|-----|-----|--|-----|-----|------|
|   |     |     |     |     |     | aaa<br>Lys        |  |     |     |     |  |     |     | 1920 |
|   |     |     |     |     |     | tca<br>Ser        |  |     |     |     |  |     |     | 1968 |
|   |     |     |     |     |     | agt<br>Ser        |  |     |     |     |  |     |     | 2016 |
| _ | _   | -   | Thr |     |     | ctg<br>Leu        |  |     |     |     |  |     |     | 2064 |
|   |     | Leu |     |     |     | gcc<br>Ala<br>695 |  |     |     |     |  |     |     | 2112 |
|   | Val |     |     |     |     | tgt<br>Cys        |  |     |     | Gln |  |     |     | 2160 |
|   |     |     |     |     | lle | gag<br>Glu        |  |     | Gln | •   |  |     | Lys | 2208 |
|   |     |     |     | Asn |     | tac<br>Tyr        |  | Asn | -   |     |  | Tyr |     | 2256 |

|     | acg<br>Thr        |     |     |     |     |     |     |     |     |     | Ser |     |     |     |     | 2304 |
|-----|-------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|------|
|     | atg<br>Met<br>770 |     |     |     |     |     |     |     |     |     |     | _   |     |     |     | 2352 |
| •   | tca<br>Ser        |     |     |     |     |     |     |     | _   | _   |     |     |     |     | _   | 2400 |
|     | atc<br>He         |     |     |     |     |     |     |     |     |     |     |     |     |     |     | 2448 |
| •   | gcc<br>Ala        |     | -   |     |     | _   |     |     |     |     |     |     |     |     |     | 2496 |
|     | aac<br>Asn        |     |     |     |     |     |     |     |     |     |     |     | _   |     |     | 2544 |
|     | ctc<br>Leu<br>850 |     |     |     |     |     | _   |     |     |     |     |     |     | _   |     | 2592 |
| _   | atc<br>He         | _   | _   |     | _   |     |     |     |     |     |     |     |     |     |     | 2640 |
|     | acc<br>Thr        |     |     |     |     |     |     |     |     |     |     |     |     |     |     | 2688 |
| gct | cac               | gtg | gag | ctg | gct | gag | agg | atg | agg | aag | cgc | gac | CCC | ggc | agt | 2736 |

| Ala His Val Glu Leu Ala Glu Arg Met Arg Lys Arg Asp Pro Gly Ser<br>900 905 910   |             |
|--|-------------|
| gcg ccc agc ctg ggt gac cga gtc ccc tat gtg atc att ggt gct gct  | 2784        |
| Ala Pro Ser Leu Gly Asp Arg Val Pro Tyr Val lle Ile Gly Ala Ala  |             |
| 915 920 925  |             |
|  |             |
| aag ggt gtg gcc gcc tac atg aag tcg gag gac ccc ctg ttt gtg ctg  | 2832        |
| Lys Gly Val Ala Ala Tyr Met Lys Ser Glu Asp Pro Leu Phe Val Leu  |             |
| 930 935 940  |             |
|  |             |
| gag cac agc ctg ccc atc gac act cag tac tac ctg gag cag cag ctg  | 2880        |
| Glu His Ser Leu Pro Ile Asp Thr Gln Tyr Tyr Leu Glu Gln Gln Leu  |             |
| 945 950 955 960  |             |
|  |             |
| gcc aag ccg ctc ttg cgc atc ttt gag ccc atc ctg ggt gag ggc cgt  | 2928        |
| Ala Lys Pro Leu Leu Arg IIe Phe Glu Pro IIe Leu Gly Glu Gly Arg  |             |
| 965 970 975  |             |
|  | 0070        |
| gca gag tot gtg ctg ctg cgc ggt gac cac aca cga tgc aag act gtg  | 2976        |
| Ala Glu Ser Val Leu Leu Arg Gly Asp His Thr Arg Cys Lys Thr Val  |             |
| 980 985 990  |             |
| ete een een een ata ann ann ete tta acc tte een een cac cac een  | 3024        |
| ctc acc agc aag gtg ggc ggc ctc ttg gcc ttc acc aag cgc cgc aac<br>Leu Thr Ser Lys Val Gly Gly Leu Leu Ala Phe Thr Lys Arg Arg Asn     | 00Z-T       |
| 995 1000 1005  |             |
| 999 1000 1000  |             |
| tgt tgc att ggc tgc cgc tcc gta atc gac cat caa gga gcc gtg  | 2060        |
| Cys Cys lie Gly Cys Arg Ser Val lie Asp His Gin Gly Ala Val  | <b>3003</b> |
|  | 3069        |
|  | 3003        |
|  | 3009        |
|  | 3114        |
| 1010 1015 1020   |             |
| 1010 1015 1020  tgt aag ttc tgt cag cca cgg gag tcg gag ctc tat cag aag gag  |             |
| tgt aag ttc tgt cag cca cgg gag tcg gag ctc tat cag aag gag Cys Lys Phe Cys Gin Pro Arg Giu Ser Giu Leu Tyr Gin Lys Giu                |             |
| tgt aag ttc tgt cag cca cgg gag tcg gag ctc tat cag aag gag Cys Lys Phe Cys Gin Pro Arg Giu Ser Giu Leu Tyr Gin Lys Giu                |             |
| tgt aag ttc tgt cag cca cgg gag tcg gag ctc tat cag aag gag Cys Lys Phe Cys Gin Pro Arg Glu Ser Glu Leu Tyr Gin Lys Glu 1025 1030 1035 | 3114        |

35

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1045 1050 1040 aca cag tgt caa cgc tgc cag ggc agc ttg cat gag gac gtc atc 3204 Thr Gln Cys Gln Arg Cys Gln Gly Ser Leu His Glu Asp Val Ile 1060 1055 3249 tgt acc agc cgt gac tgt ccc atc ttc tac atg cgc aag aag gtg Cys Thr Ser Arg Asp Cys Pro 11e Phe Tyr Met Arg Lys Lys Val 1075 1070 1080 3294 cgc aag gac ctg gaa gac cag gaa cgg ctg ctg cag cgc ttt gga Arg Lys Asp Leu Glu Asp Gln Glu Arg Leu Leu Gln Arg Phe Gly 1085 1090 1095 ccg ccc ggc cct gag gcc tgg tga 3318 Pro Pro Gly Pro Glu Ala Trp 1100 1105 <210> 89 <211> 1105 <212> PRT <213> Mus musculus <400> 89 Met Asp Cys Lys Arg Arg Gin Gly Pro Gly Pro Gly Val Pro Pro Lys 10 5 15 Arg Ala Arg Gly His Leu Trp Asp Glu Asp Glu Pro Ser Pro Ser Gln 20 25 30 Phe Giu Ala Asn Leu Ala Leu Leu Giu Giu Ile Giu Ala Giu Asn Arg

40

| Leu        | GIn<br>50  | Glu        | Ala        | Glu        | Glu        | Glu<br>55   | Leu        | Gin         | Leu         | Pro        | Pro<br>60  | Glu        | Gly        | Thr        | Val        |
|------------|------------|------------|------------|------------|------------|-------------|------------|-------------|-------------|------------|------------|------------|------------|------------|------------|
| Gly<br>65  | Gly        | GIn        | Phe        | Ser        | Thr<br>70  | Ala         | Asp        | lle         | Asp         | Pro<br>75  | Arg        | Тгр        | Arg        | Arg        | Pro<br>80  |
| Thr        | Leu        | Arg        | Ala        | Leu<br>85  | Asp        | Pro         | Ser        | Thr         | Glu<br>90   | Pro        | Leu        | He         | Phe        | GIn<br>95  | Gin        |
| Leu        | Glu        | lle        | Asp<br>100 | His        | Tyr        | Val         | Gly        | Ser<br>105  | Ala         | Pro        | Pro        | Leu        | Pro<br>110 | Glu        | Gly        |
| Pro        | Leu        | Pro<br>115 |            | Arg        | Asn        | Ser         | Val<br>120 | Pro         | lle         | Leu        | Arg        | Ala<br>125 | Phe        | Gly        | Val        |
| Thr        | Asp<br>130 |            | Gly        | Phe        | Ser        | Va I<br>135 | Cys        | Cys         | His         | lle        | GIn<br>140 |            | Phe        | Ala        | Pro        |
| Tyr<br>145 |            | Tyr        | Thr        | Pro        | Ala<br>150 |             | Pro        | Gly         | Phe         | Gly<br>155 |            | Glu        | His        | Leu        | Ser<br>160 |
| Glu        | Leu        | Gln        | Gin        | Glu<br>165 |            | Asn         | Ala        | Ala         | l le<br>170 |            | Arg        | Asp        | GIn        | Arg<br>175 |            |
| Gly        | Lys        | Glu        | Leu<br>180 |            | Gly        | Pro         | Ala        | Va l<br>185 |             | Ala        | lle        | Glu        | Leu<br>190 |            | Ser        |

| Arg         | Glu         | Ser<br>195 | Met        | Phe        | Gly         | Tyr        | His<br>200 | Gly        | His        | Gly         | Pro        | Ser<br>205 | Pro        | Phe        | Leu        |
|-------------|-------------|------------|------------|------------|-------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|
| Arg         | l le<br>210 | Thr        | Leu<br>·   | Ala        | Leu         | Pro<br>215 | Arg        | Leu        | Met        | Ala         | Pro<br>220 | Ala        | Arg        | Arg        | Leu        |
| Leu<br>225  | Glu         | Gln        | Gly        | Val        | Arg<br>230  | Val        | Pro        | Gly        | Leu        | Gly<br>235  | Thr        | Pro        | Ser        | Phe        | Ala<br>240 |
| Pro         | Tyr         | Glu        | Ala        | Asn<br>245 | Val         | Asp        | Phe        | Glu        | lle<br>250 | Arg         | Phe        | Met        | Val        | Asp<br>255 | Ala        |
| Asp         | lle         | Val        | Gly<br>260 | Cys        | Asn         | Trp        | Leu        | Glu<br>265 | Leu        | Pro         | Ala        | Gly        | Lys<br>270 | Tyr        | Val        |
| Arg         | Arg         | Ala<br>275 | Glu        | Lys        | Lys         | Ala        | Thr<br>280 | Leu        | Cys        | GIn         | Leu        | Glu<br>285 | Val        | Asp        | Val        |
| Leu<br>,    | Trp<br>290  | Ser        | Asp        | Val        | ile         | Ser<br>295 | His        | Pro        | Pro        | Glu         | Gly<br>300 | Gln        | Trp        | Gln        | Arg        |
| l le<br>305 |             | Pro        | Leu        | Arg        | Va i<br>310 | Leu        | Ser        | Phe        | Asp        | l le<br>315 |            | Cys        | Ala        | Gly        | Arg<br>320 |
| Lys         | Gly         | He         | Phe        | Pro<br>325 | Glu         | Pro        | Glu        | Arg        | Asp<br>330 | Pro         | Val        | lle        | Gln        | lle<br>335 | Cys        |
| Ser         | Leu         | Gly        | Leu        | Arg        | Trp         | Gly        | Glu        | Pro        | Glu        | Pro         | Phe        | Leu        | Arg        | Leu        | Ala        |

|            |             |            | 340        |             |            |            |            | 345        |            |             |             |            | 350        |            |            |
|------------|-------------|------------|------------|-------------|------------|------------|------------|------------|------------|-------------|-------------|------------|------------|------------|------------|
| Leu        | Thr         | Leu<br>355 | Arg        | Pro         | Cys        | Ala        | Pro<br>360 | lle        | Leu        | Gly         | Ala         | Lys<br>365 | Val        | Gln        | Ser        |
|            | Glu<br>370  | Arg        | Glu        | Glu         | Asp        | Leu<br>375 | Leu        | Gln        | Ala        | Trp         | Ala<br>380  | Asp        | Phe        | lie        | Leu        |
| Ala<br>385 | Met         | Asp        | Pro        | Asp         | Val<br>390 | lle        | Thr        | Gly        | Tyr        | Asn<br>395  | []e         | Gln        | Asn        | Phe        | A1a<br>400 |
| Leu        | Pro         | Tyr        | Leu        | l le<br>405 | Ser        | Arg        | Ala        | Gln        | Ala<br>410 | Leu         | Lys         | Vai        | Asp        | Arg<br>415 | Phe        |
| Pro        | Phe         | Leu        | Gly<br>420 | Arg         | Val        | Thr        | Gly        | Leu<br>425 | Arg        | Ser         | Asn         | lle        | Arg<br>430 | Asp        | Ser        |
| Ser        | Phe         | Gin<br>435 |            | Arg         | Gln        | Val        | Gly<br>440 | Arg        | Arg        | Asp         | Ser         | Lys<br>445 | Val        | ile        | Ser        |
| Met        | Va I<br>450 |            | Arg        | Val         | Gln        | Met<br>455 |            | Met        | Leu        | GIn         | Va I<br>460 | Leu        | Leu        | Arg        | Glu        |
| His<br>465 | Lys         | Leu        | Arg        | Ser         | Tyr<br>470 |            | Leu        | Asn        | Ala        | Va I<br>475 |             | Phe        | His        | Phe        | Leu<br>480 |
| Gly        | Glu         | Gln        | Lys        | Glu<br>485  |            | Val        | Gin        | His        | Ser<br>490 |             | He          | Thr        | Asp        | Leu<br>495 | GIn        |

| Asn         | Gly        | <b>A</b> sn | Glu<br>500       | Gin        | Thr        | Arg        | Arg        | Arg<br>505 | Leu        | Ala                | Val <sub>.</sub> | Tyr        | Cys<br>510 | Leu        | Lys        |
|-------------|------------|-------------|------------------|------------|------------|------------|------------|------------|------------|--------------------|------------------|------------|------------|------------|------------|
| Asp         | Ala        | Phe<br>515  | Leu              | Pro        | Leu        | Arg        | Leu<br>520 | Leu        | Glu        | <b>A</b> rg        | Leu              | Met<br>525 | Val        | Leu        | Vál        |
| <b>A</b> sn | Asn<br>530 | Val         | Glu              | Met        | Ala        | Arg<br>535 | Val        | Thr        | Gly        | Val                | Pro<br>540       | Leu        | Gly        | Tyr        | Leu        |
| Leu<br>545  | Thr        | Arg         | Gly              | GIn        | GIn<br>550 | Val        | Lys        | Val        | Vai        | Ser<br>555         | GIn              | Leu        | Leu        | Arg        | Gin<br>560 |
| Ala         | Met        | Arg         | Gin              | Gly<br>565 | Leu        | Leu        | Met        | Pro        | Val<br>570 | Val                | Lys              | Thr        | Glu        | Gly<br>575 | Ser        |
| Glu         | Asp        | Tyr         | Thr<br>580       | Gly        | Ala        | Thr        | Val        | lle<br>585 | Glu        | Pro                | Leu              | Lys        | Gly<br>590 | Tyr        | Tyr        |
| Asp         | Val        | Pro<br>595  | lle <sup>1</sup> | Ala        | Thr        | Leu        | Asp<br>600 | Phe        | Ser        | Ser                | Leu              | Tyr<br>605 | Pro        | Ser        | lle        |
| Met         | Met<br>610 | Ala         | His              | Asn        | Leu        | Cys<br>615 | Tyr        | Thr        | Thr        | Leu                | Leu<br>620       | Arg        | Pro        | Gly        | Ala        |
| Ala<br>625  | Gin        | Lys         | Leu              | Gly        | Leu<br>630 | Lys        | Pro        | Asp        | Glu        | Phe<br><b>63</b> 5 | lle              | Lys        | Thr        | Pro        | Thr<br>640 |

| i |            |            |                    |            |            |            |            |            |            |            |            |             |            |            |            |            |
|---|------------|------------|--------------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|
| · | Gly        | Asp        | Glu                | Phe        | Vai<br>645 | Lys        | Ser        | Ser        | Val        | Arg<br>650 | Lys        | Gly         | Leu        | Leu        | Pro<br>655 | Gin        |
|   | ile        | Leu        | Glu                | Asn<br>660 | Leu        | Leu        | Ser        | Ala        | Arg<br>665 | Lys        | Arg        | Ala         | Lys        | Ala<br>670 | Glu        | Leu        |
|   | Ala        | GIn        | Glu<br>675         | Thr        | Asp        | Pro        | Leu        | Arg<br>680 | Arg        | Gln        | Val        | Leu         | Asp<br>685 | Gly        | Arg        | Gln        |
|   | Leu        | Ala<br>690 | Leu                | Lys        | Val        | Ser        | Ala<br>695 | Asn        | Ser        | Val        | Tyr        | Gly<br>700  | Phe        | Thr        | Gly        | Ala        |
|   | GIn<br>705 | Val        | Gly                | Lys        | Leu        | Pro<br>710 | Cys        | Leu        | Glu        | lle        | Ser<br>715 | Gln         | Ser        | Val        | Thr        | Gly<br>720 |
|   | Phe        | Gly        | Arg                | GIn        | Met<br>725 | lle        | Glu        | Lys        | Thr        | Lys<br>730 | Gln        | Leu         | Val        | Glu        | Ser<br>735 | Lys        |
|   | Tyr        | Thr        | Val                | Giu<br>740 | Asn        | Gly        | Tyr        | Asp        | Ala<br>745 | Asn        | Ala        | Lys         | Val        | Val<br>750 | Tyr        | Gly        |
|   | Asp        | Thr        | <b>A</b> sp<br>755 | Ser        | Val        | Met        | Cys        | Arg<br>760 | Phe        | Gly        | Val        | Ser         | Ser<br>765 | Val        | Ala        | Glu        |
|   | Ala        | Met<br>770 | Ser                | Leu        | Gly        | Arg        | Glu<br>775 | Ala        | Ala        | Asn        | Trp        | Va I<br>780 | Ser        | Ser        | His        | Phe        |

| Pro<br>785  | Ser        | Pro        | He         | Arg        | Leu<br>790  | Glu        | Phe                | Glu        | Lys        | Va I<br>795 | Tyr        | Phe        | Pro         | Tyr         | Leu<br>800  |
|-------------|------------|------------|------------|------------|-------------|------------|--------------------|------------|------------|-------------|------------|------------|-------------|-------------|-------------|
| Leu         | He         | Ser        | Lys        | Lys<br>805 | Arg         | Tyr        | Ala                | Gly        | Leu<br>810 | Leu         | Phe        | Ser        | Ser         | Arg<br>815  | Ser         |
| Asp         | Ala        | His        | Asp<br>820 | Lys        | Met         | Asp        | Cys                | Lys<br>825 | Gly        | Leu         | Glu        | Ala        | Va I<br>830 | Arg         | Arg         |
| Asp         | Asn        | Cys<br>835 | Pro        | Leu        | Val         | Ala        | <b>A</b> sn<br>840 | Leu        | Val        | Thr         | Ser        | Ser<br>845 | Leu         | <b>A</b> rg | Arg         |
| He          | Leu<br>850 | Val        | Asp        | Arg        | <b>A</b> sp | Pro<br>855 | Asp                | Gly        | Ala        | Val         | Ala<br>860 | His        | Ala         | Lys         | Asp         |
| Va I<br>865 | He         | Ser        | Asp        | Leu        | Leu<br>870  | Cys        | Asn                | Arg        | lle        | Asp<br>875  | lle        | Ser        | Gln         | Leu         | Va I<br>880 |
| lle         | Thr        | Lys        | Glu        | Leu<br>885 | Thr         | Arg        | Ala                | Ala        | Ala<br>890 | Asp         | Tyr        | Ala        | Gly         | Lys<br>895  | Gin         |
| Ala         | His        | Val        | Glu<br>900 |            | Ala         | Glu        | Arg                | Met<br>905 | Arg        | Lys         | Arg        | Asp        | Pro<br>910  | Gly         | Ser         |
| Ala         | Pro        | Ser<br>915 |            | Gly        | Asp         | Arg        | Va I<br>920        |            | Tyr        | Val         | lle        | lle<br>925 |             | Ala         | Ala         |

Lys Gly Val Ala Ala Tyr Met Lys Ser Glu Asp Pro Leu Phe Val Leu

|     | 930                        | 935                        | 940                    |
|-----|----------------------------|----------------------------|------------------------|
| Glu | His Ser Leu Pro lle        |                            | Leu Glu Gln Gln Leu    |
| 945 | 950                        |                            | 960                    |
| Ala | Lys Pro Leu Leu Arg        | : lie Phe Glu Pro lie      | e Leu Gly Glu Gly Arg  |
|     | 965                        | 970                        | 975                    |
| Ala | Glu Ser Val Leu Leu<br>980 | Arg Gly Asp His Thr<br>985 | Arg Cys Lys Thr Val    |
| Leu | Thr Ser Lys Val Gly        | Gly Leu Leu Ala Ph         | ne Thr Lys Arg Arg Asn |
|     | 995                        | 1000                       | 1005                   |
| Cys | Cys lle Gly Cys Ar         | rg Ser Val IIe Asp H       | lis Gin Gly Ala Val    |
|     | 1010                       | 1015                       | 1020                   |
| Cys | Lys Phe Cys Gin Pr         | ro Arg Glu Ser Glu L       | .eu Tyr Gin Lys Giu    |
|     | 1025                       | 1030                       | 1035                   |
| Val | Ser His Leu Asn Al         | a Leu Glu Glu Arg F        | Phe Ser Arg Leu Trp    |
|     | 1040                       | 1045                       | 1050                   |
| Thr | Gin Cys Gin Arg Cy         | vs Gin Gly Ser Leu H       | dis Glu Asp Val IIe    |
|     | 1055                       | 1060                       | 1065                   |
| Cys | Thr Ser Arg Asp Cy         | vs Pro lle Phe Tyr N       | let Arg Lys Lys Val    |

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|--|---|---|----------------|
| ttcaccaccg gaacattgag                                | -,<br>acaaaatoto gacggtgacg ct                                  | tgaggttga aaaaaccaat 18                       | 30             |
| cgaaccgcag acgtaccagg                                | aaccgaacca tgtatcaacg co  | cattgaaga agaagaagaa 24                       | <del>1</del> 0 |
| gaagaaggtg aaaaacgaaa                                | gattgagaat ttgtttgctt tg  | gagcaacca aacctcagga 30                       | 00             |
| aaaaagagtt aaggtgggag                                | tgtctggttc aaccggttta ta  | atceggtte aaattaaacc 36                       | <b>30</b>      |
| tottacagtt aaccgggttt                                | tgtgtttggt tcgattgttc at  | taaaagaaa gaagactott 42                       | 20             |
| gtcgtcgatt agtgccaaag                                | ttgaaagttg aaaccttttc to  | cagaattit ctgctcagtt 48                       | 30             |
| ctgagttttt ttttcccgcc                                | atg gaa atc gac tcc gag<br>Met Glu Ile Asp Ser Glu<br>1 5       |   | 33             |
| aag caa toc gat tac a<br>Lys Gin Ser Asp Tyr A<br>15 | at tog ctg gtacgaactc tat<br>sn Ser Leu                         | ttacttta togacttgta 58                        | 37             |
| gtgaaagaca aatgtaatca                                | ttcgtggtgg tgactgtttc ta  | acttataag tgtacgggct 64                       | 47             |
| agggtttgtt atctgattct                                | gagtttttgc aattgaagca g   | gat gag aga ttc gag 70<br>Asp Glu Arg Phe Glu | 03             |
| •  |   | 20  |                |
| lle Gin Lys Giu Met T                                | ac aga ggt cag caa tac ag<br>yr Arg Gly Gln Gln Tyr Se<br>30 35 | 20<br>gt cag att tac ttt 75                   | 51             |

| tgg aaa tot cat ttg cot g gtoagtgott ttgtttotot catatttagc Trp Lys Ser His Leu Pro 60   | 848  |
|---|------|
| acaacaacga agagcagttt ttgagaattt tcttgggtta gatataatta ggtgaaatca   | 908  |
| gtgattitta gggattittg ctatcttatg gattacagtt gagaaagatt gctagtattg   | 968  |
| tttaaattat agatotgaat gigaattica tittigoag ig tigt aag git tig<br>Val Cys Lys Val Leu<br>65   | 1021 |
| gga ctt gaa aaa gga aaa gaa tgc ata att gtg gga acc ttg ttc aaa<br>Gly Leu Glu Lys Gly Lys Glu Cys IIe IIe Val Gly Thr Leu Phe Lys<br>70 75 80  | 1069 |
| cac atg aag ctt aaa cct tgt gtt ctc gat gaa tat tct aaa gag<br>His Met Lys Leu Lys Pro Cys Val Leu Asp Glu Tyr Ser Lys Glu<br>85 90 95          | 1114 |
| gttggttttt attaacctct actgtttttt tgagctatgt ctatgctgaa tcaatctgag   | 1174 |
| tatatttaac ataatgcag agg tca gtt act ccg ctt gtt aaa cca cat aac Arg Ser Val Thr Pro Leu Val Lys Pro His Asn 100 105                            | 1226 |
| ttt atg cat cct gat gat aat ctg atc ctc gaa gac gag agt ggg aga Phe Met His Pro Asp Asp Asn Leu IIe Leu Glu Asp Glu Ser Gly Arg 110 115 120 125 | 1274 |
| gtt aag ott got ggt toe goa ott toa oot gog att tat gtg aca g<br>Val Lys Leu Ala Gly Ser Ala Leu Ser Pro Ala IIe Tyr Val Thr<br>130 135 140     | 1320 |
| gtattgcaaa tgggttctta ctgtttttac tgtatgattt tttccttctt tacaatgtgg   | 1380 |
| caaatcttag agattttgat caagctttcc totottaaaa gatgggttct ttaagaaaat   | 1440 |

| taacgttgaa gcctcccgtg cattgtag gt gtt gtt gtt gca ctg cat ggg<br>Gly Val Val Ala Leu His Gly<br>145   | 1491 |
|---|------|
| aag gaa act aat gct ggt gaa ttc ttt gtt gag gat gta cta gaa gct<br>Lys Glu Thr Asn Ala Gly Glu Phe Phe Val Glu Asp Val Leu Glu Ala<br>150 155 160 | 1539 |
| ggt tta cca cct cag att gag cgg cct atc gat cta c gtaagtctag  Gly Leu Pro Pro Gin Ile Glu Arg Pro Ile Asp Leu  165 170 175                        | 1586 |
| ctatgttctc ttccttttgc taacctcatg gctcaatcat ttctataagc aatctctcat   | 1646 |
| gatacatcca tattgcatct gcag ag gaa gat aaa tat gtc gtg tta ttg Gin Giu Asp Lys Tyr Va! Vai Leu Leu 180 185   | 1696 |
| tog ggc ctt tgt att gga agc aaa tog gct aat ccc ctg cag ttt cag<br>Ser Gly Leu Cys IIe Gly Ser Lys Ser Ala Asn Pro Leu Gln Phe Gln<br>190 195 200 | 1744 |
| ctt ctt gtt gac cat ata act ggg cat ctc gga gat gag gag<br>Leu Leu Val Asp His lle Thr Gly His Leu Gly Asp Glu Glu<br>205 210 215                 | 1786 |
| gttcaaatct cttaacttgc aggttgttca acatatttct ttccttaatt tatactttat   | 1846 |
| ggtttgaaca g gaa caa ggc ctt gca gca cag ata gtt cat gta gta att<br>Glu Gln Gly Leu Ala Ala Gln Ile Val His Val Val Ile<br>220 225                | 1896 |
| gct gga aac tot ttt gaa ttt ccc cgc aaa ctc att aat ggc cag<br>Ala Gly Asn Ser Phe Glu Phe Pro Arg Lys Leu lle Asn Gly Gin<br>230 235. 240        | 1941 |

| gtacttataa cttttgttgc tgatatattc tcagatacag ttccagtaat tatctgcccc   | 2001 |
|---|------|
| agttatgtet tatgatettt attggttgat etttgtag aac ttg gee teg aaa gat<br>Asn Leu Ala Ser Lys Asp<br>245   | 2057 |
| caa tog aca ctg tat gag coc atc aaa gag ctt gat atc atg tta agc<br>Gin Ser Thr Leu Tyr Giu Pro iie Lys Giu Leu Asp iie Met Leu Ser<br>250 255 260 265 | 2105 |
| cag gtcagttaac tggatctacg tgtgtgttat cgatatctat tgagatgaaa<br>Gln   | 2158 |
| gttcaaactc ctgtttttt ttttgtggat tgtttttag ata gct gca gga gtt<br>lle Ala Ala Gly Val<br>270   | 2212 |
| tca gta gat atc atg cca ggc acg aat gat cca gct aac ttc gca ttg Ser Val Asp IIe Met Pro Gly Thr Asn Asp Pro Ala Asn Phe Ala Leu 275 280 285           | 2260 |
| cct cag cag gtctgcaaat acataagaaa cattcaaaat cccgcatttt Pro Gln Gln 290   | 2309 |
| gtatogataa ototgattoa taggoootto tottitgtto ag oot otg aat aga<br>Pro Leu Asn Arg   | 2363 |
| tgt ctt ttc cct gga tct tca cct tat aac acc ttc aga tca tgt aca<br>Cys Leu Phe Pro Gly Ser Ser Pro Tyr Asn Thr Phe Arg Ser Cys Thr<br>295 300 305 310 | 2411 |
| aat cct cac tca ttt gat gtc gat aat atc ag gtatgattat tattaatagt Asn Pro His Ser Phe Asp Val Asp Asn Ile Arg 315 320                                  | 2463 |

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| tgaatacaat ctctctgatt ttacaacgat aaaattcttg ggtttatctg actgaaaacc   | 2523         |
|---|--------------|
| teatatgggg gcattttgca g a ttt ctt gga act tct ggt cag aac atc gat  Phe Leu Gly Thr Ser Gly Gln Asn Ile Asp  325 330   | 2575         |
| gac ctt ggc aag tac tca gag gct aag agc aag ctt gat ttt gtg gaa<br>Asp Leu Gly Lys Tyr Ser Glu Ala Lys Ser Lys Leu Asp Phe Val Glu<br>335 340 345   | 2623         |
| aga acg ctg agg tgg aga cat ctt gcc cca act gca cct aat aca ctc g Arg Thr Leu Arg Trp Arg His Leu Ala Pro Thr Ala Pro Asn Thr Leu 350 355 360   | 2672         |
| gtaagaatte teettgeest geaagattas ttttttgaas taageesata aaaaaatgat   | 2732         |
| cctttgagtt ctatttggtt ttgattcact tgcgtacag gt tgt tat cct ttc Gly Cys Tyr Pro Phe 365   | 2785         |
|   |              |
| acc gat aga gac cct ttc ttg att gaa acc tgc ccg cat gtc tac ttc Thr Asp Arg Asp Pro Phe Leu IIe Glu Thr Cys Pro His Val Tyr Phe 370 380   | 2833         |
| Thr Asp Arg Asp Pro Phe Leu IIe Glu Thr Cys Pro His Val Tyr Phe   | 2833<br>2883 |
| Thr Asp Arg Asp Pro Phe Leu IIe Glu Thr Cys Pro His Val Tyr Phe 370 375 380  gtc ggg aat caa gat aaa tat gac aac cgt ttg ata aag g gtaaaagcac Va! Gly Asn Gln Asp Lys Tyr Asp Asn Arg Leu IIe Lys             |              |
| Thr Asp Arg Asp Pro Phe Leu IIe Glu Thr Cys Pro His Val Tyr Phe 370 375 380  gtc ggg aat caa gat aaa tat gac aac cgt ttg ata aag g gtaaaagcac Va! Gly Asn Gln Asp Lys Tyr Asp Asn Arg Leu IIe Lys 385 390 395 | 2883         |

| 410          |            | 415         |             | 420                         |            |      |
|--------------|------------|-------------|-------------|-----------------------------|------------|------|
| aatttgagca g | gaatttgaga | ccatttaccc  | tcatagattg  | cagattotaa                  | atctcaaaat | 3102 |
| caccatgtet a |            |             |             | ggaa tgt ca<br>ı Glu Cys Hi |            | 3154 |
| agc ttt agc  | act cag at | a aac caa t | ca taacatte | gag ttgctact                | tt         | 3201 |
| Ser Phe Ser  |            |             |             | ,                           |            |      |
|              | 435        | 4           | 140         |                             |            |      |
| ggtagattat   | ttcctgtctt | gaagatgtaa  | tgttgagctt  | tttcagtaac                  | acactcctat | 3261 |
| gttctaacca a | aatgtttgtt | aaaaatcctt  | tttcttgagt  | ggaacttcca                  | aatctttgga | 3321 |
| tatattggta : | atgotoattg | ttttgtccta  | attttctaaa  | aatotogaca                  | cgagttctta | 3381 |
| ggtagtcaca · | taaaggacaa | aaagggccga  | ccagatagtg  | togtggtogt                  | tggtcagaag | 3441 |
| aacgtgaaaa ( | gactgcaaaa | ataatottaa  | aaaaagcaac  | aagtgcacag                  | aatctcatgc | 3501 |
| aaatgtotot ( | ctctctcttc | tcaacggcta  | tatccatcca  | cacttattac                  | attataaaat | 3561 |
| taattaaatg ( | caataatgta | acgcattata  | ttctccaacg  | gtccattttc                  | ccgcatttcc | 362  |
| ctaacctttc ( | ctttataacg | caaaacagtt  | tcatcttcta  | cacttaacac                  | tttaatcctc | 368  |
| tct          |            |             |             |                             |            | 3684 |

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**<400> 91** 

| Met<br>1  | Glu        | lle        | Asp        | Ser<br>5  | Giu         | Lys        | lle        | His        | Glu<br>10 | Arg       | Lys        | GIn        | Ser        | Asp<br>15 | Tyr       |
|-----------|------------|------------|------------|-----------|-------------|------------|------------|------------|-----------|-----------|------------|------------|------------|-----------|-----------|
| Asn       | Ser        | Leu        | Asp<br>20  | Ğlu       | <b>A</b> rg | Phe        | Glu        | l le<br>25 | Gin       | Lys       | Glu        | Met        | Tyr<br>30  | Arg       | Gly       |
| Gin       | GIn        | Tyr<br>35  | Ser        | GIn       | He          | Tyr        | Phe<br>40  | Ala        | Arg       | Leu       | His        | Leu<br>45  | Met        | Arg       | Thr       |
| Leu       | Leu<br>50  | Tyr        | Ser        | Leu       | Ala         | Pro<br>55  | Thr        | Trp        | Lys       | Ser       | His<br>60  | Leu        | Pro        | Val       | Cys       |
| Lys<br>65 | Val        | Leu        | Gly        | Leu       | Glu<br>70   | Lys        | Gly        | Lys        | Glu       | Cys<br>75 | lle        | lle        | Val        | Gly       | Thr<br>80 |
| Leu       | Phe        | Lys        | His        | Met<br>85 | Lys         | Leu        | Lys        | Pro        | Cys<br>90 | Val       | Leu        | Asp        | Glu        | Tyr<br>95 | Ser       |
| Lys       | Glu        | Arg        | Ser<br>100 |           | Thr         | Pro        | Leu        | Vai<br>105 | Lys       | Pro       | His        | Asn        | Phe<br>110 | Met       | His       |
| Pro       | Asp        | Asp<br>115 | Asn        | Leu       | lle         | Leu        | Glu<br>120 | Asp        | Glu       | Ser       | Gly        | Arg<br>125 |            | Lys       | Leu       |
| Ala       | Gly<br>130 |            | Ala        | Leu       | Ser         | Pro<br>135 | Ala        | lle        | Tyr       | Val       | Thr<br>140 | Gly        | Val        | Val       | Val       |

| Ala<br>145  | Leu        | His         | Gly        | Lys        | Glu<br>150 | Thr        | Asn        | Ala        | Gly         | Glu<br>155 | Phe        | Phe        | Val        | Glu        | Asp<br>160 |
|-------------|------------|-------------|------------|------------|------------|------------|------------|------------|-------------|------------|------------|------------|------------|------------|------------|
| Val         | Leu        | Glu         | Ala        | Gly<br>165 | Leu        | Pro        | Pro        | Gln        | l le<br>170 | Glu        | Arg        | Pro        | lle        | Asp<br>175 | Leu        |
| Gin         | Glu        | Asp         | Lys<br>180 | Tyr        | Val        | Val        | Leu        | Leu<br>185 | Ser         | Gly        | Leu        | Cys        | lle<br>190 | Gly        | Ser        |
| Lys         | Ser        | Ala<br>195  | Asn        | Pro        | Leu        | Gin        | Phe<br>200 | Gin        | Leu         | Leu        | Val        | Asp<br>205 | His        | He         | Thr        |
| Gly         | His<br>210 | Leu         | Gly        | Asp        | Glu        | Glu<br>215 | Glu        | Gin        | Gly         | Leu        | Ala<br>220 | Ala        | GIn        | He         | Val        |
| His<br>225  | Val        | Val         | He         | Ala        | Gly<br>230 | Asn        | Ser        | Phe        | Glu         | Phe 235    | Pro        | Arg        | Lys        | Leu        | 11e<br>240 |
| <b>A</b> sn | Gly        | Gln         | Asn        | Leu<br>245 |            | Ser        | Lys        | Asp        | GIn<br>250  | Ser        | Thr        | Leu        | Tyr        | Glu<br>255 | Pro        |
| He          | Lys        | Glu         | Leu<br>260 |            | lle        | Met        | Leu        | Ser<br>265 | Gin         | lle        | Ąlа        | Ala        | Gly<br>270 | Val        | Ser        |
| Val         | Asp        | l le<br>275 |            | Pro        | Gly        | Thr        | Asn<br>280 |            | Pro         | Ala        | Asn        | Phe<br>285 | Ala        | Leu        | Pro        |

Gin Gin Pro Leu Asn Arg Cys Leu Phe Pro Gly Ser Ser Pro Tyr Asn 290 295 300

Thr Phe Arg Ser Cys Thr Asn Pro His Ser Phe Asp Val Asp Asn Ile 305 310 315 320

Arg Phe Leu Gly Thr Ser Gly Gln Asn I le Asp Asp Leu Gly Lys Tyr 325 330 335

Ser Glu Ala Lys Ser Lys Leu Asp Phe Val Glu Arg Thr Leu Arg Trp 340 345 350

Arg His Leu Ala Pro Thr Ala Pro Asn Thr Leu Gly Cys Tyr Pro Phe 355 360 365

Thr Asp Arg Asp Pro Phe Leu IIe Glu Thr Cys Pro His Val Tyr Phe 370 375 380

Val Gly Asn Gln Asp Lys Tyr Asp Asn Arg Leu lle Lys Gly Ser Glu 385 390 395 400 .

Gly Gln Leu Val Arg Leu lle Cys lle Pro Lys Phe Cys Glu Thr Gly 405 410 415

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Phe Ser Thr Gln Ile Asn Gin Ser

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| ⟨222⟩  | (2198) (2269)    |            |            |            |            |     |
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|        | (2352) (2443)    |            |            |            |            |     |
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| ⟨221⟩  | CDS              | •          |            |            |            |     |
|        | (2545) (2672)    |            |            |            |            |     |
|        | (                |            |            |            |            |     |
| <220>  |                  |            |            |            |            |     |
| ⟨221⟩  | CDS              |            |            |            |            |     |
|        | (2772) (2873)    |            |            |            |            |     |
|        |                  |            |            |            |            |     |
| ⟨220⟩  |                  |            |            |            |            |     |
| ⟨221⟩  | CDS              |            |            |            |            |     |
|        | (2965) (3032)    |            |            |            |            |     |
|        |                  |            |            |            |            |     |
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| ⟨221⟩  | CDS              |            |            |            |            |     |
|        | (3122) (3181)    |            |            |            |            |     |
|        | ••               |            |            |            |            |     |
| <400>  | 92               |            |            |            |            |     |
|        | tcct tagcagaaag  | gcgctccatt | tctctggcgt | aaaccaaagg | agatccttga | 60  |
| -0-0   |                  |            |            | •          |            |     |
| acteti | tect geaceattge  | tcttaaaacc | cttctccggc | acgaattctt | ccaaccctgc | 120 |
|        | <b>3</b>         |            |            | _          |            |     |
| ttcacc | caccg gaacattgag | acaaaatctc | gacggtgacg | ctgaggttga | aaaaaccaat | 180 |
| 20000  |                  |            |            |            |            |     |
|        |                  |            | tatataaaa  | oosttraaga | araaraaraa | 240 |

| gaagaaggtg aaaaacgaaa gattgagaat ttgtttgctt tgagcaacca aacctcagga   | 300 |
|---|-----|
| aaaaagagtt aaggtgggag tgtctggttc aaccggttta tatccggttc aaattaaacc   | 360 |
| tottacagtt aaccgggttt tgtgtttggt tcgattgttc ataaaagaaa gaagactctt   | 420 |
| gtcgtcgatt agtgccaaag ttgaaagttg aaaccttttc tcagaatttt ctgctcagtt   | 480 |
| ctgagttttt ttttcccgcc atg gaa atc gac tcc gag aaa att cac gaa agg Met Glu lle Asp Ser Glu Lys lle His Glu Arg 1 5 10                              | 533 |
| aag caa too gat tac aat tog otg gtacgaacto tattacttta togacttgta  | 587 |
| Lys Gin Ser Asp Tyr Asn Ser Leu 15  |     |
| gtgaaagaca aatgtaatca ttcgtggtgg tgactgtttc tacttataag tgtacgggct   | 647 |
| agggtttgtt atctgattct gagtttttgc aattgaagca g gat gag aga ttc gag Asp Glu Arg Phe Glu 20  | 703 |
| ata cag aag gag atg tac aga ggt cag caa tac agt cag att tac ttt<br>lle Gin Lys Glu Met Tyr Arg Gly Gin Gin Tyr Ser Gin ile Tyr Phe<br>25 30 35 40 | 751 |
| get egt ett eat ete atg aga aca ett ete tae tet ett get eet act Ala Arg Leu His Leu Met Arg Thr Leu Leu Tyr Ser Leu Ala Pro Thr 45 50 55          | 799 |
| tgg aaa tot cat ttg cot g gtoagtgott ttgtttotot catatttago Trp Lys Ser His Leu Pro 60   | 848 |
| acaacaacga agagcagttt ttgagaattt tcttgggtta gatataatta ggtgaaatca   | 908 |
| gtgattttta gggatttttg ctatcttatg gattacagtt gagaaagatt gctagtattg   | 968 |

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| tttaaattat agatotgaat gtgaatttoa tttttgcag tg tgt aag gtt Val Cys Lys Val 65   |   |  |  |  |  |  |  |  |  |  |
|--|---|--|--|--|--|--|--|--|--|--|
| gga ctt gaa aaa gga aaa gaa tgc ata att gtg gga acc ttg ttc<br>Gly Leu Glu Lys Gly Lys Glu Cys Ile Ile Val Gly Thr Leu Phe<br>70 75 80   | •   |  |  |  |  |  |  |  |  |  |
| cac atg aag ctt aaa cct tgt gtt ctc gat gaa tat tct aaa gag<br>His Met Lys Leu Lys Pro Cys Val Leu Asp Glu Tyr Ser Lys Glu   |   |  |  |  |  |  |  |  |  |  |
| 85 90 95   |   |  |  |  |  |  |  |  |  |  |
| gttggttttt attaacctct actgtttttt tgagctatgt ctatgctgaa tcaa  | tctgag 1174                                     |  |  |  |  |  |  |  |  |  |
| tatatttaac ataatgcag agg tca gtt act ccg ctt gtt aaa cca cat aac Arg Ser Val Thr Pro Leu Val Lys Pro His Asn 100 105   |   |  |  |  |  |  |  |  |  |  |
|  |   |  |  |  |  |  |  |  |  |  |
| ttt atg cat cct gat gat aat ctg atc ctc gaa gac gag agt ggg Phe Met His Pro Asp Asp Asn Leu IIe Leu Glu Asp Glu Ser Gly 110 115 120  |   |  |  |  |  |  |  |  |  |  |
| Phe Met His Pro Asp Asp Asn Leu IIe Leu Glu Asp Glu Ser Gly  | Arg<br>125<br>g 1320                            |  |  |  |  |  |  |  |  |  |
| Phe Met His Pro Asp Asp Asn Leu IIe Leu Glu Asp Glu Ser Gly 110 115 120  gtt aag ctt gct ggt tcc gca ctt tca cct gcg att tat gtg aca Val Lys Leu Ala Gly Ser Ala Leu Ser Pro Ala IIe Tyr Val Thr   | Arg<br>125<br>g 1320                            |  |  |  |  |  |  |  |  |  |
| Phe Met His Pro Asp Asp Asp Leu IIe Leu Glu Asp Glu Ser Gly 110 115 120  gtt aag ctt gct ggt tcc gca ctt tca cct gcg att tat gtg aca Val Lys Leu Ala Gly Ser Ala Leu Ser Pro Ala IIe Tyr Val Thr 130 135 140   | Arg<br>125<br>g 1320<br>atgtgg 1380             |  |  |  |  |  |  |  |  |  |
| Phe Met His Pro Asp Asp Asp Leu IIe Leu Glu Asp Glu Ser Gly 110 115 120  gtt aag ctt gct ggt tcc gca ctt tca cct gcg att tat gtg aca Val Lys Leu Ala Gly Ser Ala Leu Ser Pro Ala IIe Tyr Val Thr 130 135 140  gtattgcaaa tgggttctta ctgtttttac tgtatgattt tttccttctt tacaa | Arg 125 g 1320 atgtgg 1380 gaaaat 1440 ggg 1491 |  |  |  |  |  |  |  |  |  |

| Lys Glu Thr Asn Ala Gly Glu Phe Phe Val Glu Asp Val Leu Glu Ala<br>150 155 160  |      |
|---|------|
| ggt tta cca cct cag att gag cgg cct atc gat cta c gtaagtctag  Gly Leu Pro Pro Gln Ile Glu Arg Pro Ile Asp Leu  165 170 175                        | 1586 |
| ctatgttctc ttccttttgc taacctcatg gctcaatcat ttctataagc aatctctcat   | 1646 |
| gatacatcca tattgcatct gcag ag gaa gat aaa tat gtc gtg tta ttg Gin Glu Asp Lys Tyr Val Val Leu Leu 180 185   | 1696 |
| tcg ggc ctt tgt att gga agc aaa tcg gct aat ccc ctg cag ttt cag<br>Ser Gly Leu Cys IIe Gly Ser Lys Ser Ala Asn Pro Leu Gln Phe Gln<br>190 195 200 | 1744 |
| ctt ctt gtt gac cat ata act ggg cat ctc gga gat gag gag<br>Leu Leu Val Asp His lle Thr Gly His Leu Gly Asp Glu Glu<br>205 210 215                 | 1786 |
| gttcaaatct cttaacttgc aggttgttca acatatttct ttccttaatt tatactttat   | 1846 |
| ggtttgaaca g gaa caa ggc ctt gca gca cag ata gtt cat gta gta att Glu Gln Gly Leu Ala Ala Gln Ile Val His Val Val Ile 220 225                      | 1896 |
| gct gga aac tot ttt gaa ttt ccc cgc aaa ctc att aat ggc cag<br>Ala Gly Asn Ser Phe Glu Phe Pro Arg Lys Leu lle Asn Gly Gln<br>230 235 240         | 1941 |
| gtacttataa cttttgttgc tgatatattc tcagatacag ttccagtaat tatctgcccc   | 2001 |
| agttatgtct tatgatcttt attggttgat ctttgtag aac ttg gcc tcg aaa gat Asn Leu Ala Ser Lys Asp 245   | 2057 |

| caa tog aca otg tat gag occ atc aaa gag ott gat atc atg tta ago<br>Gin Ser Thr Leu Tyr Giu Pro lie Lys Giu Leu Asp lie Met Leu Ser<br>250 255 260 265 | 2105 |
|---|------|
| cag gtcagttaac tggatctacg tgtgtgttat cgatatctat tgagatgaaa<br>Gin   | 2158 |
| gttcaaactc ctgtttttt ttttgtggat tgtttttag ata gct gca gga gtt<br>ile Ala Ala Gly Val<br>270   | 2212 |
| tca gta gat atc atg cca ggc acg aat gat cca gct aac ttc gca ttg<br>Ser Val Asp lie Met Pro Gly Thr Asn Asp Pro Ala Asn Phe Ala Leu<br>275 280 285     | 2260 |
| cct cag cag gtotgcaaat acataagaaa cattcaaaat cccgcatttt<br>Pro Gin Gin<br>290   | 2309 |
| gtatogataa ototgatica taggocotto tottttgtto ag oot otg aat aga<br>Pro Leu Asn Arg   | 2363 |
| tgt ctt ttc cct gga tct tca cct tat aac acc ttc aga tca tgt aca<br>Cys Leu Phe Pro Gly Ser Ser Pro Tyr Asn Thr Phe Arg Ser Cys Thr<br>295 300 305 310 | 2411 |
| aat oot oac toa tit got git gat aat atc ag gitatgattat tattaatagi<br>Asn Pro His Ser Phe Ala Val Asp Asn Ile Arg<br>315 320                           | 2463 |
| tgaatacaat ctctctgatt ttacaacgat aaaattcttg ggtttatctg actgaaaacc   | 2523 |
| tcatatgggg gcattltgca g a ttt ctt gga act tct ggt cag aac atc gat  Phe Leu Gly Thr Ser Gly Gln Asn ile Asp  325 330                                   | 2575 |

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| gac ctt ggc aag tac tca gag                                | gct aag agc aag ctt   | gat ttt gtg gaa 2623    |
|--|-----------------------|-------------------------|
| Asp Leu Gly Lys Tyr Ser Glu                                | Ala Lys Ser Lys Leu   | Asp Phe Val Glu         |
| 335  | 340                   | 345                     |
|  |                       |                         |
| aga acg ctg agg tgg aga cat                                | ctt gcc cca act gca   | cct aat aca ctc g 2672  |
| Arg Thr Leu Arg Trp Arg His                                | Leu Ala Pro Thr Ala   | Pro Asn Thr Leu         |
| 350  | 355                   | 360                     |
| · .  |                       | •                       |
| gtaagaatto toottgooot gcaaga                               | attac tititigaac taa  | goccata aaaaaatgat      |
|  |                       | _                       |
| cctttgagtt ctatttggtt ttgat                                | toact tecetacae et    | tgt tat cct ttc 2785    |
|  |                       | Cys Tyr Pro Phe         |
|  | -                     | 365                     |
|  |                       |                         |
| acc gat aga gac cct ttc ttg                                | att gaa acc tgc ccg   | cat gtc tac ttc 2833    |
| Thr Asp Arg Asp Pro Phe Leu                                |                       |                         |
| 370 375  | 380                   | ino tai iyi ino         |
| 370 073  | 300                   |                         |
| ato and est oss ast see tot                                | gae age out the ata   | aag g gtaaaagcac 2883   |
| gtc ggg aat caa gat aaa tat<br>Val Gly Asp Glo Asp Lys Tyr |                       |                         |
| Val Gly Asn Gln Asp Lys Tyr<br>385                         |                       | Lys                     |
| 385 390  | 395                   |                         |
|  |                       |                         |
| cttacacaga gattagaaat aacat                                | terer frigicaaac atea | aggettt aacttttett 2943 |
| and the state of the same                                  |                       |                         |
| gggtaaatat gaatgctgca g gg                                 |                       |                         |
| Gly  | Ser Glu Gly Gln Leu   |                         |
|  | 400                   | 405                     |
|  |                       |                         |
| tgc att cct aag ttc tgt gag                                | <del>-</del>          |                         |
| Cys lle Pro Lys Phe Cys Glu                                | Thr Gly Ile Ala Val   | Ala                     |
| 410  | 415                   | 420                     |
|  |                       |                         |
| aatttgagca gaatttgaga ccatt                                | taccc tcatagattg cag  | attotaa atotoaaaat 3102 |
|  |                       |                         |
| caccatgtct atttcgcag gtg aac                               | cta aga aat ctg gaa   | a tgt cac act tta 3154  |
| Val Ası  | n Leu Arg Asn Leu Glu | ı Cys His Thr Leu       |
|  |                       |                         |

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#### 227/236

425 430

age ttt age act cag ata aac caa tea taacattgag ttgetaettt 3201 Ser Phe Ser Thr Gln Ile Asn Gln Ser 435

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<211> 440

<212> PRT

<213> Arabidopsis thaliana

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|---|------------|------------|------------|------------|-----------|------------|------------|------------|------------|-----------|------------|------------|------------|------------|-----------|------------|
|   | Gln        | Gln        | Tyr<br>35  | Ser        | Gin       | He         | Tyr        | Phe<br>40  | Ala        | Arg       | Leu        | His        | Leu<br>45  | Met        | Arg       | Thr        |
|   | Leu        | Leu<br>50  | Tyr        | Ser        | Leu       | Ala        | Pro<br>55  | Thr        | Trp        | Lys       | Ser        | His<br>60  | Leu        | Pro        | Val       | Cys        |
|   | Lys<br>65  | Val        | Leu        | Gly        | Leu       | Glu<br>70  | Lys        | Gly        | Lys        | Glu       | Cys<br>75  | He         | He         | Val        | Gly       | Thr<br>80  |
|   | Leu        | Phe        | Lys        | His        | Met<br>85 | Lys        | Leu        | Lys        | Pro        | Cys<br>90 | Val        | Leu        | Asp        | Glu        | Tyr<br>95 | Ser        |
| • | Lys        | Glu        | Arg        | Ser<br>100 | Val       | Thr        | Pro        | Leu        | Val<br>105 | Lys       | Pro        | His        | Asn        | Phe<br>110 | Met       | His        |
|   | Pro        | Asp        | Asp<br>115 | Asn        | Leu       | He         | Leu        | Glu<br>120 | Asp        | Glu       | Ser        | Gly        | Arg<br>125 | Val        | Lys       | Leu        |
|   | Ala        | Gly<br>130 | Ser        | Ala        | Leu       | Ser        | Pro<br>135 | Ala        | He         | Tyr       | Val        | Thr<br>140 | Gly        | Val        | Val       | Val        |
|   | Ala<br>145 | Leu        | His        | Gly        | Lys       | Glu<br>150 | Thr        | Asn        | Ala        | Gly       | Glu<br>155 | Phe        | Phe        | Val        | Glu       | Asp<br>160 |
|   | Val        | Leu        | Glu        | Ala        | Gly       | Leu        | Pro        | Pro        | Gin        | lle       | Glu        | Arg        | Pro        | He         | Asp       | Leu        |

|                    |             |            |            | 165        |            |            |            |            | 170        |            |            |            |             | 175        |             |
|--------------------|-------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|-------------|------------|-------------|
| Gin                | Glu         | Asp        | Lys<br>180 | Tyr        | Val        | Val        | Leu        | Leu<br>185 | Ser        | Gly        | Leu        | Cys        | l le<br>190 | Gly        | Ser         |
| Lys                | Ser         | Ala<br>195 | Asn        | Pro        | Leu        | GIn        | Phe<br>200 | Gin        | Leu        | Leu        | Val        | Asp<br>205 | His         | He         | Thr         |
| Gly i              | His<br>210  | Leu        | Gly        | Asp        | Glu        | Glu<br>215 | Glu        | Gin        | Gly        | Leu        | Ala<br>220 | Ala        | GIn         | He         | Val         |
| His '              | Val         | Val        | He         | Ala        | Gly<br>230 | Asn        | Ser        | Phe        | Glu        | Phe<br>235 | Pro        | Arg        | Lys         | Leu        | lle<br>240  |
| Asn (              | Gly         | Gln        | Asn        | Leu<br>245 | Ala        | Ser        | Lys        | Asp        | Gln<br>250 | Ser        | Thr        | Leu        | Tyr         | Glu<br>255 | Pro         |
| lle                | Lys         | Glu        | Leu<br>260 | Asp        | lle        | Met        | Leu        | Ser<br>265 | Gln        | lle        | Ala        | Ala        | Gly.<br>270 | Val        | Ser         |
| Val <sub>,</sub> i | <b>A</b> sp | 11e<br>275 |            | Pro        | Gly        | Thr        | Asn<br>280 | Asp        | Pro        | Ala        | Asn        | Phe<br>285 | Ala         | Leu        | Pro         |
| Gin (              | GIn<br>290  | Pro        | Leu        | Asn        | Arg        | Cys<br>295 | Leu        | Phe        | Pro        | Gly        | Ser<br>300 | Ser        | Pro         | Tyr        | <b>As</b> n |
| Thr 1              | Phe         | Arg        | Ser        | Cys        | Thr<br>310 | Asn        | Pro        | His        | Ser        | Phe<br>315 | Ala        | Val        | Asp         | Asn        | 11e<br>320  |

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<212> DNA

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|-------------|------------|------------|------------|------------|------------|------------|-------------|-------------|------------|------------|------------|------------|------------|------------|------------|
| Ser         | Glu        | Ala        | Lys<br>340 | Ser        | Lys        | Leu        | Asp         | Phe<br>345  | Val        | Glu        | Arg        | Thr        | Leu<br>350 | Arg        | Trp        |
| Arg         | His        | Leu<br>355 | Ala        | Pro        | Thr        | Alą        | Pro<br>360  | Asn         | Thr        | Leu        | Gly        | Cys<br>365 | Tyr        | Pro        | Phe        |
| Thr         | Asp<br>370 | Arg        | Asp        | Pro        | Phe        | Leu<br>375 | He          | Glu         | Thr        | Cys        | Pro<br>380 | His        | Val        | Tyr        | Phe        |
| Va I<br>385 | Gly        | Asn        | Gln        | Asp        | Lys<br>390 | Tyr        | <b>A</b> sp | <b>A</b> sn | Arg        | Leu<br>395 | lle        | Lys        | Gly        | Ser        | Glu<br>400 |
| Gly         | Gln        | Leu        | Val        | Arg<br>405 | Leu        | lle        | Cys         | lle         | Pro<br>410 | Lys        | Phe        | Cys        | Glu        | Thr<br>415 | Gly        |
| ile         | Ala        | Val        | Ala<br>420 | Val        | Asn        | Leu        | Arg         | Asn<br>425  | Leu        | Glu        | Cys        | His        | Thr<br>430 | Leu        | Ser        |
| Phe         | Ser        | Thr<br>435 | Gln        | lle        | <b>Asn</b> | Gin        | Ser<br>440  |             |            |            | -          |            |            |            |            |

WO 2004/087960 PCT/JP2004/004378

#### 231/236

<213> Mus musculus

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| ttctaatttc          | ccaactctaa | ccccaagete | tccgttttac | tgtttagtat | tctaggctgg | 240 |
| cagtttgagt          | ctgtaccagg | caaaaaacgt | tccaaatcaa | gatagacagg | atggagaacc | 300 |
| aatcacagag          | ctggatttcc | tttcaaattc | taccaatggc | tattgtgcag | gagactttga | 360 |
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420

454

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| 540 <sub>.</sub> | ctgttctggg | tggcgttccc | tttttggccc | tcactccacc | ccctacccgc | accccactc  |
| 600              | ctaggccatc | tgatggccga | ctctttccag | caatgggcct | tttgtgtgtc | gcatataaag |
| 660              | cataatgttg | ctggttagtt | ctccggggta | gagtcaagag | tatgcagcta | ttttgataca |
| 720              | gctcctccat | actttctcta | ctccttgggt | atccctttag | agggttgcag | atccacctat |
| 780              | gtttgctagg | ccacttctgt | ctgtgggcat | cattagctga | gtgatccatc | tgggagccct |
| 840              | atcttgctag | tttcgataaa | atctgggtcc | agacagotac | gtctcacaag | cccggcata  |
| 900              | gataaggcag | tggatccctg | gattatgggg | tttggatgct | ggtgtcagcg | tgtatgcaat |
| 960              | ctccttccaa | gtctctgtaa | tccaaacttt | tcatctcagc | gtccatcctt | tctctacatg |
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| 1080             | ggttttgggc | gggtatccta | cttatatctt | gaaattgtat | atgtgtttag | catgagtttc |
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| 1320             | tgctatgaac | taaataaggc | ctggctatta | tttccagctt | atctgggttc | gttgagggg  |
| 1380             | gcccaggaga | ctggatatat | ggggcatctc | cttaccagtt | atgtgtcctt | atagtggago |
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